Attorney Docket No.: 018501-000710US

Date of Deposit December 6, 2001

### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

MURRAY et al.

Application No.: Not yet assigned

Filed: Herewith

For: NOVEL METHODS OF DIAGNOSIS OF ANGIOGENESIS, COMPOSITIONS AND METHODS OF SCREENING FOR ANGIOGENESIS MODULATORS Examiner:

Not yet assigned

Art Unit:

PRELIMINARY AMENDMENT FOR PRIORITY CLAIM

Assistant Commissioner for Patents Washington, D.C. 20231

Sir:

Prior to examination of the above-referenced application, please enter the following amendments and remarks.

#### **IN THE SPECIFICATION:**

Please delete the paragraph on page 1, lines 7-11/

Please add the following paragraph on page 1, following line 6, the CROSS-REFERENCES TO RELATED APPLICATIONS:

This application is a continuation of co-pending U.S. Application No. 09/784,356, filed February 14, 2001; and is also a continuation-in-part (CIP) of co-pending U.S. Application No. 09/637,977, filed August 11, 2000, which claims benefit of

Application No.: 09/784,356

Page 2

priority of U.S. Application No. 60/148,425, filed August 11, 1999. All of the applications are incorporated herein by reference.

Although a marked-up version of the deleted and inserted paragraphs does not appear to be required (see, 37 C.F.R. § 1.121(b)(iii)), for convenience a marked up copy of the deleted paragraph that indicates the changes set forth in the inserted paragraph is attached hereto.

If the Examiner believes a telephone conference would expedite prosecution of this application, please telephone the undersigned at 415-576-0200.

Respectfully submitted,

Jean M. Lockyer, Ph.D. Reg. No. 44,879

TOWNSEND and TOWNSEND and CREW LLP Two Embarcadero Center,  $8^{\rm th}$  Floor San Francisco, California 94111-3834

Tel: (415) 576-0200 Fax: (415) 576-0300

**JML** SF 1296905 v1

Application No.: 09/784,356

Page 3

# **VERSION WITH MARKINGS TO SHOW CHANGES MADE**

[The present] This application is a continuation of co-pending U.S.

Application No. 09/784,356, filed February 14, 2001; and is also a continuation-in-part

(CIP) of co-pending [United States Patent Application "Novel Methods Of Diagnosis Of Angiogenesis, Compositions And Methods Of Screening For Angiogenesis Modulators",

Attorney Docket No. A65110-1] U.S. Application No. 09/637,977, filed [on] August 11, 2000, which claims [the] benefit of priority to [U.S.S.N.] U.S. Application No. 60/148,425 filed August 11, 1999.[, both of which] All of the applications are incorporated herein by reference.

SF 1296905 v1

Attorney Docket No.: 018501-000710US

Date of Deposit December 6, 2001

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

MURRAY et al.

Application No.: not assigned

Filed: herewith

For: NOVEL METHODS OF DIAGNOSIS OF ANGIOGENESIS, COMPOSITIONS AND METHODS OF SCREENING FOR ANGIOGENESIS MODULATORS Examiner:

Not yet assigned

Art Unit:

Not assigned

**COMMUNICATION UNDER** 

37 C.F.R. §§ 1.821-1.825

AND

PRELIMINARY AMENDMENT

Box SEQUENCE Assistant Commissioner for Patents Washington, D.C. 20231

Sir:

Please amend the application as set forth below.

The computer readable form of the "Sequence Listing" in the current application is identical to that filed in the parent application, Application No. 09/784,356, filed February 14, 2001. In accordance with 37 C.F.R. § 1.821(e), please use the computer readable form filed in that application as the computer readable for for the instant application. It is understood that the Patent and Trademark Office will make the necessary change in application number and filing date for the computer readable form that will be used for the instant application. A paper copy of the "Sequence Listing" filed in the parent application is included in the instant amendment for incorporation into the specification.

The information contained in the computer readable disk submitted in the parent application was prepared through the use of the software program "FastSEQ" and is identical to the paper copy.

Application No.: 09/784,356

Page 2

### In the Specification:

Please replace the paragraph beginning at page 42, line 23, with the following:

--Various tag polypeptides and their respective antibodies are well known in the art. Examples include poly-histidine (poly-his) or poly-histidine-glycine (poly-his-gly) tags; HIS6 (SEQ ID NO:135) and metal chelation tags, the flu HA tag polypeptide and its antibody 12CA5 [Field et al., Mol. Cell. Biol., 8:2159-2165 (1988)]; the c-myc tag and the 8F9, 3C7, 6E10, G4, B7 and 9E10 antibodies thereto [Evan et al., Molecular and Cellular Biology, 5:3610-3616 (1985)]; and the Herpes Simplex virus glycoprotein D (gD) tag and its antibody [Paborsky et al., Protein Engineering, 3(6):547-553 (1990)]. Other tag polypeptides include the Flag-peptide [Hopp et al., BioTechnology, 6:1204-1210 (1988)]; the KT3 epitope peptide [Martin et al., Science, 255:192-194 (1992)]; tubulin epitope peptide [Skinner et al., J. Biol. Chem., 266:15163-15166 (1991)]; and the T7 gene 10 protein peptide tag [Lutz-Freyermuth et al., Proc. Natl. Acad. Sci. USA, 87:6393-6397 (1990)].--

Please replace the paragraph beginning at page 82, line 3, with the following:

```
--AAA4 DNA sequence (SEQ ID NO:1):
Gene name: CGI-100 protein
Unigene number: Hs.275253
Probeset Accession #: AA089688
Nucleic Acid Accession #: NM_016040 cluster
Coding sequence: 142-831 (predicted start/stop codons underlined)--
```

Please replace the paragraph beginning at page 82, line 32, with the following:

```
--AAA7 DNA sequence (SEQ ID NO:2):

Gene name: Endothelial differentiation, sphingolipid G-protein-coupled receptor, 1 (EDG1)

Unigene number: Hs.154210

Probeset Accession #: M31210

Nucleic Acid Accession #: NM_001400 cluster

Coding sequence: 251-1396 (predicted start/stop codons underlined)--
```

Please replace the paragraph beginning at page 83, line 20, with the following:

Application No.: 09/784,356

Page 3

--AAB3 DNA sequence (SEQ ID NO:3):

Gene name: Solute carrier family 20 (phosphate transporter), member 1, Human

leukaemia virus receptor 1 (GLVR1)

Unigene number: Hs.78452 Probeset Accession #: L20859

Nucleic Acid Accession #: NM\_005415 cluster

Coding sequence: predicted 371-2410 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 84, line 16, with the following:

--AAB4 DNA sequence (SEQ ID NO:4):

Gene name: Matrix metalloproteinase 10 (stromelysin 2)

Unigene number: Hs.2258

Probeset Accession #: X07820

Nucleic Acid Accession #: NM\_002425

Coding sequence: predicted 23-1453 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 84, line 55, with the following:

--AAB6 DNA sequence (SEQ ID NO:5):

Gene name: Podocalyxin-like Unigene number: Hs.16426 Probeset Accession #: U97519

Nucleic Acid Accession #: NM\_005397 cluster

Coding sequence: 251-1837 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 86, line 26, with the following:

--AAB8 DNA sequence (SEQ ID NO:6):

Gene name: EGF-containing fibulin-like extracellular matrix protein 1

Unigene number: Hs.76224 Probeset Accession #: U03877

Nucleic Acid Accession #: NM\_004105 Transcript variant 1

Coding sequence: 150-1631 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 87, line 13, with the following:

--AAB9 DNA sequence (SEQ ID NO:7):

Gene name: Melanoma adhesion molecule, MUC 18 glycoprotein

Unigene number: Hs.211579 Probeset Accession #: M28882

Nucleic Acid Accession #: NM\_006500 cluster

MURRAY *et al.* Application No.: 09/784,356 Page 4

Coding sequence: 27-1967 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 5

Please replace the paragraph beginning at page 88, line 14, with the following:

--AAC1 DNA sequence (SEQ ID NO:8):

Gene name: Matrix metalloproteinase 1 (interstitial collagenase)

Unigene number: Hs.83169 Probeset Accession #: X54925

Nucleic Acid Accession #: NM\_002421 cluster

Coding sequence: 69-1478 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 88, line 56, with the following:

--AAC3 DNA sequence (SEQ ID NO:9):

Gene name: Branched chain aminotransferase 1, cytosolic

Unigene number: Hs.157205

Probeset Accession #: AA423987

Nucleic Acid Accession #: NM\_005504 cluster

Coding sequence: 1-1155 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 89, line 17, with the following:

--ACG4 DNA sequence SEQ ID NO:10):

Gene name: Pentaxin-related gene, rapidly induced by IL-1 beta

Unigene number: Hs.2050

Probeset Accession #: M31166

Nucleic Acid Accession #: NM\_002852 cluster

Coding sequence: 68-1213 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 89, line 57, with the following:

--ACK5 DNA sequence (SEQ ID NO:11):

Gene name: Von Willebrand factor; Coagulation factor VIII

Unigene number: Hs.110802 Probeset Accession #: M10321

Nucleic Acid Accession #: NM 000552

Coding sequence: 311-8752 (predicted start/stop codons underlined) --

Application No.: 09/784,356

Page 6

Please replace the paragraph beginning at page 92, line 11, with the following:

--AAC7 DNA sequence (SEQ ID NO:12):

Gene name: KIAA1294 protein
Probeset Accession #: AA432248
Nucleic Acid Accession #: AB037715

Coding sequence: 370-3489 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 93, line 65, with the following:

--ACG8 DNA sequence (SEQ ID NO:13):

Gene name: ubiquitin E3 ligase SMURF2 Unigene number: Hs.21806 (3'UTR only)

Probeset Accession #: AA398243

Nucleic Acid Accession #: AF301463 cluster

Coding sequence: 9-2255 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 94, line 44, with the following:

--ACH1 DNA sequence (SEQ ID NO:14):

Gene name: EST

Unigene number: Hs.30089

Probeset Accession #: AA410480

CAT cluster#: 96816\_1

Coding sequence: Partial sequence, possible frameshift. Predicted stop codon

underlined .--

Please replace the paragraph beginning at page 95, line 2, with the following:

--ACJ2 DNA sequence (SEQ ID NO:15):

Gene name: Complement component Clq receptor

Unigene number: Hs.97199

Probeset Accession #: AA487558 Nucleic Acid Accession #: NM\_012072

Coding sequence: 149-2107. Predicted start/stop codons underlined--

Application No.: 09/784,356

Page 7

Please replace the paragraph beginning at page 96, line 55, with the following:

--ACJ3 DNA sequence (SEQ ID NO:16):

Gene name: FLT1/vascular endothelial growth factor receptor

Unigene number: Hs.138671

Probeset Accession #: AA047437 Nucleic Acid Accession #: NM\_002019

Coding sequence: 250-4266 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 98, line 56, with the following:

--ACJ9 DNA sequence (SEQ ID NO:17):

Gene name: Purine nucleoside phosphorylase

Unigene number: Hs.75514 Probeset Accession #: K02574

Nucleic acid Accession #: X00737 cluster

Coding sequence: 110-979 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 99, line 21, with the following:

--ACK4 DNA sequence (SEQ ID NOS:18 and 19):

Gene name: EST

Unigene number: Hs.265499 Probeset Accession #: R68763 CAT cluster#: Cluster 46668\_2

Sequence: Both the EST corresponding to the probeset accession and exon prediction; number and the CAT cluster align with the Homo sapiens BAC clone AC009414 RP11-490M8. Using FGENESH, 2 exons predicted on this BAC clone upstream of the probeset.

Predicted exon 1: bases 5808-5837 of BAC clone AC009414--

Please replace the paragraph beginning at page 99, line 58, with the following:

--Predicted ACK4 gene seq (predicted start/stop codons underlined) (SEQ ID NO:19):--

Application No.: 09/784,356

Page 8

Please replace the paragraph beginning at page 100, line 20, with the following:

--AAA8 DNA sequence (SEQ ID NO:20):

Gene name: ETL protein, with extended open reading frame

Unigene number: Hs.57958
Probeset Accession #: D58024
Nucleotide Accession #: AF192403

Coding sequence: 151-2136. Underlined sequences correspond to extended

sequence not included in AF192403.--

Please replace the paragraph beginning at page 101, line 11, with the following:

--AAC6 DNA sequence (SEQ ID NO:21):

Gene name: Homo sapiens cDNA FLJ13465 fis, clone PLACE1003493, weakly similar

to endothelial cell multimerin precursor

Unigene number: Hs.134797 Probeset Accession #: AA025351 Nucleotide Accession #: AK023527

Nucleotide Accession #: AKU23527 Coding sequence: predicted 75-2921

Extended sequence: 729-3465 (underlined sequence)--

Please replace the paragraph beginning at page 102, line 18, with the following:

--ACH7 DNA sequence (SEQ ID NOS:22-24):

Gene name: ESTs

Uniquene number: Hs.3807

Probeset Accession #: AA292694

BAC Accession #: AL161751

FGENESH predicted exons: FGENESH predicts 2 exons on the minus strand of

AL161751 upstream of the ACH7 probeset .--

Please replace the paragraph beginning at page 102, line 26, with the following:

--FGENESH predicted exon 1 (SEQ ID NO:22):--

Please replace the paragraph beginning at page 102, line 32, with the following:

--FGENESH predicted exon 2 (SEQ ID NO:23):--

Application No.: 09/784,356

Page 9

Please replace the paragraph beginning at page 102, line 63, with the following:

--ACH7 predicted coding seq (predicted start/stop codons underlined) (SEQ ID NO:24):--

Please replace the paragraph beginning at page 103, line 31, with the following:

--AAD3 DNA sequence (SEQ ID NO:25):

Gene name: ESTs

Unigene number: Hs.17404 Probeset Accession #: N39584 Nucleic Acid Accession #: N39584

Coding sequence: no identified ORF; possible frameshifts--

Please replace the paragraph beginning at page 103, line 64, with the following:

--AAD4 DNA sequence (SEQ ID NO:26):

Gene name: ERG

Unigene number: Hs.279477 / Hs.45514

Probeset Accession #: R32894 Nucleic Acid Accession #: M17254

Coding sequence: 257-1645 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 104, line 58, with the following:

--AAD5 DNA sequence (SEQ ID NO:27):

Gene name: activin A receptor type II-like 1 (ALK-1)

Unigene number: Hs.8881 / Hs.172670

Probeset Accession #: T57112

Nucleic Acid Accession #: NM\_000020

Coding sequence: 283-1794 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 106, line 3, with the following:

--AAD8 DNA sequence (SEQ ID NO:28):

Gene name: ESTs

Unigene number: Hs.144953

Probeset Accession #: AA404418 Nucleic Acid Accession #: n/a

Coding sequence: no ORF identified; possible frameshifts--

Application No.: 09/784,356

Page 10

Please replace the paragraph beginning at page 106, line 39, with the following:

--ACA2 DNA sequence (SEQ ID NO:29):

EST Gene name:

Unigene number: Hs.16450

Probeset Accession #: AA478778 Nucleic Acid Accession #: AA478778

Coding sequence: no ORF identified; possible frameshifts--

Please replace the paragraph beginning at page 107, line 1, with the following:

--ACA4 DNA sequence (SEQ ID NO:30):

Gene name: alpha satellite junction DNA sequence

Unigene number: Hs.247946 Probeset Accession #: M21305 Nucleic Acid Accession #: M21305

Coding sequence: 1-165 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 107, line 13, with the following:

--ACG6 DNA sequence (SEQ ID NO:31):

Gene name: intercellular adhesion molecule 2 (ICAM2)

Unigene number: Hs.83733 Probeset Accession #: M32334

Nucleic Acid Accession #: NM\_000873

Coding sequence: 63-890 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 107, line 40, with the following:

--ACG7 DNA sequence (SEQ ID NO:32):

Gene name: Cadherin 5, VE-cadherin (CDH5)

Unigene number: Hs.76206 Probeset Accession #: X79981

Nucleic Acid Accession #: NM\_001795

Coding sequence: 25-2379 (predicted start/stop codons underlined)--

Davit Agassii signaja 1800

Ţ,

MURRAY et al.

Application No.: 09/784,356

Page 11

Please replace the paragraph beginning at page 108, line 48, with the following:

--ACG9 DNA sequence (SEQ ID NO:33):

Gene name: lysyl oxidase-like 2 (LOXL2)

Unigene number: Hs.83354 Probeset Accession #: U89942

Nucleic Acid Accession #: NM\_002318 cluster

Coding sequence: 248-2572 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 109, line 47, with the following:

--ACH2 DNA sequence (SEQ ID NO:34):

Gene name: TIE tyrosine-protein kinase

Unigene number: Hs.78824 Probeset Accession #: X60957

Nucleic Acid Accession #: NM\_005424 cluster

Coding sequence: 37-3452 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 110, line 53, with the following:

--ACH3 DNA sequence (SEQ ID NO:35):

Gene name: placental growth factor (PGF; PlGF1; VEGF-related protein)

Unigene number: Hs.2894

Probeset Accession #: X54936

Nucleic Acid Accession #: NM 002632 cluster

Coding sequence: 322-768 (predicted start/stop codons underlined) --

Please replace the paragraph beginning at page 111, line 22, with the following:

--ACH4 DNA sequence (SEQ ID NO:36):

Gene name: nidogen 2 (NID2) Unigene number: Hs.82733 Probeset Accession #: D86425

Nucleic Acid Accession #: NM\_007361 cluster

Coding sequence: 1-4131 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 12

Please replace the paragraph beginning at page 112, line 44, with the following:

--ACH5 DNA sequence (SEQ ID NO:37):

Gene name: SNL (singed-like; sea urchin fascin homolog-like)

Unigene number: Hs.118400 Probeset Accession #: U03057

Nucleic Acid Accession #: NM\_003088

Coding sequence: 112-1593 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 113, line 32, with the following:

--ACH6 DNA sequence (SEQ ID NO:38):

Gene name: endothelial protein C receptor (EPCR; PROCR)

Unigene number: Hs.82353 Probeset Accession #: L35545

Nucleic Acid Accession #: NM\_006404

Coding sequence: 25-741 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 113, line 63, with the following:

--ACH8 DNA sequence (SEQ ID NO:39):

Gene name: melanoma adhesion molecule (MCAM; MUC18)

Unigene number: Hs.211579 Probeset Accession #: D51069

Nucleic Acid Accession #: NM\_006500

Coding sequence: 27-1967 (predicted start and stop codons underlined)--

Please replace the paragraph beginning at page 114, line 64, with the following:

--ACH9 DNA sequence (SEQ ID NO:40):

Gene name: endothelin-1 (EDN1)

Unigene number: Hs.2271

Probeset Accession #: J05008

Nucleic Acid Accession #: NM\_001955

Coding sequence: 337-975 (predicted start/stop codons underlined)--

i c

MURRAY et al.

Application No.: 09/784,356

Page 13

Please replace the paragraph beginning at page 115, line 26, with the following:

--ACJ1 DNA sequence (SEQ ID NO:41):

Gene name: BMX non-receptor tyrosine kinase

Unigene number: Hs.27372 Probeset Accession #: X83107

Nucleic Acid Accession #: NM\_001721

Coding sequence: 34-2061 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 116, line 8, with the following:

--ACJ4 DNA sequence (SEQ ID NO:42):

Gene name: prostaglandin G/H synthase 2 (COX-2; PGHS-2)

Unigene number: Hs.196384 Probeset Accession #: D28235

Nucleic Acid Accession #: NM\_000963

Coding sequence: 135-1949 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 117, line 24, with the following:

--ACJ6 DNA sequence (SEQ ID NO:43):

Gene name: SEC14-like-1 Unigene number: Hs.75232 Probeset Accession #: D67029

Nucleic Acid Accession #: NM\_003003

Coding sequence: 304-2451 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 118, line 56, with the following:

--ACJ8 DNA sequence (SEQ ID NO:44):

Gene name: intercellular adhesion molecule 1 (ICAM1; CD54)

Unigene number: Hs.168383 Probeset Accession #: M24283

Nucleic Acid Accession #: NM\_000201

Coding sequence: 58-1656 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 14

Please replace the paragraph beginning at page 119, line 47, with the following:

--ACK3 DNA sequence (SEQ ID NO:45):

Gene name: angiopoietin 1 receptor (TIE-2; TEK)

Unigene number: Hs.89640 Probeset Accession #: L06139

Nucleic Acid Accession #: NM\_000459

Coding sequence: 149-3523 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 120, line 57, with the following:

--PZA6 DNA sequence (SEQ ID NO:46):

Gene name: prostate differentiation factor (PLAB; MIC-1)

Unigene number: Hs.116577

Probeset Accession #: AB000584

Nucleic Acid Accession #: NM\_004864

Coding sequence: 26-952 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 121, line 19, with the following:

--AAC8 DNA sequence (SEQ ID NO:47):

Gene name: none

Unigene number: Hs.6682

Probeset Accession #: AA227926 Nucleic Acid Accession #: none

Coding sequence: no ORF identified, possible frameshifts--

Please replace the paragraph beginning at page 122, line 12, with the following:

--AAD2 DNA sequence (SEQ ID NO:48):

Gene name: Thrombospondin-1

Unigene number: Hs.87409

Probeset Accession #: AA232645 Nucleic Acid Accession #: NM\_003246

Coding sequence: 112-3624 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 15

Please replace the paragraph beginning at page 123, line 49, with the following:

--AAD9 DNA sequence (SEQ ID NO:49):

Gene name: LIM homeobox protein cofactor (CLIM-1)

Unigene number: Hs.4980

Probeset Accession #: F13782

Nucleic Acid Accession #: AF047337

Coding sequence: 110-1231(predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 124, line 29, with the following:

--AAE1 DNA sequence (SEQ ID NO:50):

Gene name: guanine nucleotide binding protein 11

Unigene number: Hs.83381 Probeset Accession #: U31384

Nucleic Acid Accession #: NM\_004126.1

Coding sequence: 108-329 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 124, line 49, with the following:

--AAE2 DNA sequence (SEQ ID NO:51):

Gene name: Transcription factor 4 (immunoglobulin transcription factor 2)(ITF-

2) (SL3-3 Enhancer factor 2) (SEF-2)

Uniquene number: Hs.289068 Probeset Accession #: M74719

Nucleic Acid Accession #: NM\_003199.1

Coding sequence: 200-2203 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 125, line 33, with the following:

--AAE4 DNA sequence (SEQ ID NO:52):

Gene name: phosphatidylcholine 2-acylhydrolase

Unigene number: Hs.211587 Probeset Accession #: M68874 Nucleic Acid Accession #: M68874

Coding sequence: 139-2388 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 16

Please replace the paragraph beginning at page 126, line 22, with the following:

--ACA1 DNA sequence (SEQ ID NO:53):

Gene name: tissue factor pathway inhibitor 2 TFPI2, placental protein 5 (PP5)

Unigene number: Hs.78045 Probeset Accession #: D29992

Nucleic Acid Accession #: D29992.1

Coding sequence: 57-764 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 126, line 51, with the following:

--ACB8 DNA sequence (SEQ ID NO:54):

Gene name: myosin X

Unigene number: Hs.61638 Probeset Accession #: N77151

Nucleic Acid Accession #: NM\_012334

Coding sequence: 223-6399 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 128, line 54, with the following:

--ACC3 DNA sequence (SEQ ID NO:55):

Gene name: calcitonin receptor-like (CALCRL)

Unigene number: Hs.152175 Probeset Accession #: L76380

Nucleic Acid Accession #: NM\_005795

Coding sequence: 555-1940 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 129, line 46, with the following:

--ACC4 DNA sequence (SEQ ID NO:56):

Gene name: Homo sapiens mRNA; cDNA DKFZp586E1624

Unigene number: Hs.94030

Probeset Accession #: AA452000

Nucleic Acid Accession #: AL110152.1

Coding sequence: no ORF identified, possible frameshifts-

Application No.: 09/784,356

Page 17

Please replace the paragraph beginning at page 130, line 10, with the following:

--ACC5 DNA sequence (SEQ ID NO:57):

Selectin E (endothelial adhesion molecule 1) Gene name:

Unigene number: Hs.89546 Probeset Accession #: M24736

Nucleic Acid Accession #: NM\_000450

Coding sequence: 117-1949 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 131, line 15, with the following:

--ACC8 DNA sequence (SEQ ID NO:58):

Gene name: Chemokine (C-X-C motif), receptor 4 (fusin)

Unigene number: Hs.89414 Probeset Accession #: L06797

Nucleic Acid Accession #: NM\_003467

Coding sequence: 89-1147 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 131, line 52, with the following:

--ACF2 DNA sequence (SEQ ID NO:59):

Gene name: Endothelial cell-specific molecule 1

Unigene number: Hs.41716 Probeset Accession #: X89426

Nucleic Acid Accession #: NM\_007036

Coding sequence: 56-610 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 132, line 27, with the following:

--ACF4 DNA sequence (SEQ ID NO:60):

Gene name: P53-responsive gene 2 similar to D.melanogaster peroxidasin(U11052)

Unigene number: Hs.118893 Probeset Accession #: D86983 Nucleic Acid Accession #: D86983

Coding sequence: 1-4491 (predicted stop codon underlined, sequence is open at

5' end) --

Application No.: 09/784,356

Page 18

Please replace the paragraph beginning at page 133, line 61, with the following:

--ACF5 DNA sequence (SEQ ID NO:61):

Gene name: Mitogen-activated protein kinase kinase kinase kinase 4

Unigene number: Hs.3628

Probeset Accession #: N54067

Nucleic Acid Accession #: NM\_004834

Coding sequence: 80-3577 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 134, line 67, with the following:

--ACF8 DNA sequence (SEQ ID NO:62):

Gene name: Phospholipase A2, group IVC (cytosolic, calcium-independent)

Unigene number: Hs.18858

Probeset Accession #: AA054087

Nucleic Acid Accession #: NM\_003706

Coding sequence: 310-1935 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 135, line 50, with the following:

--ACG1 DNA sequence (SEQ ID NO:63):

Gene name: Carbohydrate (chondroitin 6/keratan) sulfotransferase 1

Unigene number: Hs.104576

Probeset Accession #: AA868063

Nucleic Acid Accession #: NM 003654

Coding sequence: 367-1602 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 136, line 32, with the following:

--ACG5 DNA sequence (SEQ ID NO:64):

Gene name: Multimerin
Unigene number: Hs.268107
Probeset Accession #: U27109

Nucleic Acid Accession #: U27109.1

Coding sequence: 72-3758 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 19

Please replace the paragraph beginning at page 137, line 43, with the following:

--ACC6 DNA sequence (SEQ ID NO:65):

Gene name: Homo sapiens cDNA FLJ11502 fis, clone HEMBA1002102, weakly similar

to ANKRYIN

Unigene number: Hs.213194 Probeset Accession #: AA187101 Nucleic Acid Accession #: AK021564

Coding sequence: 1-450 (predicted stop codon underlined, 5'end sequence is

open) --

Please replace the paragraph beginning at page 138, line 13, with the following:

--ACC7 DNA sequence (SEQ ID NO:66):

Gene name: Human RAL A gene Unigene number: Hs.6906

Probeset Accession #: AA083572

Nucleic Acid Accession #: contig of X15014.1 and AK026850

Coding sequence: 1-621 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 138, line 65, with the following:

--ACC9 DNA sequence (SEQ ID NO:67):

Gene name: KIAA0955 protein Unigene number: Hs.10031

Probeset Accession #: AA027168 Nucleic Acid Accession #: AB023172

Coding sequence: 314-1609 (predicted start/stop codons underlined)--

Please replace the paragraph beginning at page 140, line 23, with the following:

--ACF6 DNA sequence (SEQ ID NO:68):

Gene name: Homo sapiens cDNA FLJ10669 fis, clone NT2RP2006275, weakly similar

to Microtubule-associated protein 1B [CONTAINS: LIGHT CHAIN LC1]

Unigene number: Hs.66048

Probeset Accession #: AA609717 Nucleic Acid Accession #: AK001531

Coding sequence: 176-2194 (predicted start/stop codons underlined)--

Application No.: 09/784,356

Page 20

domains .--

Please replace the paragraph beginning at page 141, line 4, with the following:

--AAA4 Protein sequence (SEQ ID NO:69):

Gene name: CGI-100 protein

Unigene number: Hs.275253

Probeset Accession #: AA089688

Protein Accession #: NP\_057124

Signal sequence: predicted 1-23 (first underlined sequence)

Transmembrane Domain: predicted 201-217 (second underlined sequence)

emp24/gp25L/p24 domain: predicted 13-227

Summary: gp25L/emp24/p24 protein family members of the cis-Golgi network bind both COP I and II coatomer. Members of this family are implicated in bringing cargo forward from the ER and binding to coat proteins by their cytoplasmic

Please replace the paragraph beginning at page 141, line 22, with the following:

--AAA7 Protein sequence (SEQ ID NO:70):

Gene name: Endothelial differentiation, sphingolipid G-protein-coupled receptor, 1 (EDG1)

Unigene number: Hs.154210

Probeset Accession #: M31210

Protein Accession#: NP\_001391

7 Transmembrane Domains: predicted 50-71, 92-110, 122-140, 160-177, 201-222, 251-269, 281-301 (underlined sequences)

Summary: Endothelial differentiation, sphingolipid G-protein-coupled receptor, 1 may regulate the differentiation of endothelial cells. It binds the sphingolipid metabolite, sphingosine-1-phosphate, which may function as a second messenger in cell proliferation and survival.--

Please replace the paragraph beginning at page 141, line 44, with the following:

```
--AAB3 Protein sequence (SEQ ID NO:71):
Gene name: Solute carrier family 20 (phosphate transporter), member 1, Human leukaemia virus receptor 1 (GLVR1)
Unigene number: Hs.78452
Probeset Accession #: L20859
Protein Accession #: NP_005406
Transmembrane domains: predicted 24-40, 62-78, 164-180, 198-214, 232-248, 513-529, 562-578, 604-620, 655-671
Cellular Localization: Likely a Type IIIa membrane protein (Ncyt Cexo)--
```

Application No.: 09/784,356

Page 21

Please replace the paragraph beginning at page 141, line 68, with the following:

--AAB4 Protein sequence (SEQ ID NO:72):

Gene name: Matrix metalloproteinase 10 (stromelysin 2)

Unigene number: Hs.2258

Probeset Accession #: X07820 Protein Accession #: NP\_002416

Signal sequence: predicted 1-17 (underlined sequence)

Cellular Localization: predicted secreted--

Please replace the paragraph beginning at page 142, line 18, with the following:

--AAB6 Protein sequence (SEQ ID NO:73):

Gene name: Podocalyxin-like
Unigene number: Hs.16426
Probeset Accession #: U97519
Protein Accession #: NP\_005388

Transmembrane domain: predicted 432-448 (underlined sequence)
Cellular Localization: predicted Type Ia membrane protein (Nexo)--

Please replace the paragraph beginning at page 142, line 37, with the following:

--AAB8 Protein sequence (SEQ ID NO:74):

Gene name: EGF-containing fibulin-like extracellular matrix protein 1

Unigene number: Hs.76224 Probeset Accession #: U03877

Protein Accession #: NP 004096 Variant 1

Signal sequence: predicted 1-17 (underlined sequence)

Summary: This gene spans approximately 18 kb of genomic DNA and consists of 12 exons. Two transcripts with distinct 5' UTR have been described; the resulting proteins have distinct N-terminal amino acid sequences. Translation initiation from internal methionine residues was observed with in vitro translation. A signal peptide sequence is predicted for translation initiation sites 1, 2, and 4. The protein isoforms contain 5 or 6 calcium-binding EGF2 domains and 5 or 6 EGF2 domains. Mutations in this gene cause the retinal disease Malattia Leventinese. Transcript Variant: This variant (1) has a distinct 5' UTR and N-terminal protein sequence as compared to variant 2.--

Application No.: 09/784,356

Page 22

Please replace the paragraph beginning at page 142, line 64, with the following:

--AAB9 Protein sequence (SEQ ID NO:75):

Gene name: Melanoma adhesion molecule, MUC 18 glycoprotein

Unigene number: Hs.211579 Probeset Accession #: M28882 Protein Accession #: NP\_006491

Signal sequence: predicted 1-17 (first underlined sequence)

Transmembrane domain: predicted 559-575 (second underlined sequence)
Cellular localization: predicted Type Ia membrane protein (Nexo)--

Please replace the paragraph beginning at page 143, line 18, with the following:

--AAC1 Protein sequence (SEQ ID NO:76):

Gene name: Matrix metalloproteinase 1 (interstitial collagenase)

Unigene number: Hs.83169
Probeset Accession #: X54925
Protein Accession #: NP\_002412

Signal sequence: predicted 1-19 (underlined sequence) Cellular localization: predicted secreted protein--

Please replace the paragraph beginning at page 143, line 36, with the following:

--AAC3 Protein sequence (SEQ ID NO:77):

Gene name: Branched chain aminotransferase 1, cytosolic

Uniquene number: Hs.157205

Probeset Accession #: AA423987
Protein Accession #: NP\_005495
Cellular Localization: cytolasmic

Summary: The lack of the cytosolic enzyme branched-chain amino acid transaminase (BCT) causes cell growth inhibition. There may be at least 2 different clinical disorders due to a defect of branched-chain amino acid transamination: hypervalinemia and hyperleucine-isoleucinemia. Since there are 2 distinct BCATs, mitochondrial and cytosolic, it is possible that one is mutant in each of these 2 conditions.--

MURRAY et al. Application No.: 09/784,356

Page 23

Please replace the paragraph beginning at page 143, line 58, with the following:

--ACG4 Protein sequence (SEQ ID NO:78):
Gene name: Pentaxin-related gene, rapidly induced by IL-1 beta
Unigene number: Hs.2050
Probeset Accession #: M31166
Protein Accession #: NP\_002843
Signal sequence: predicted 1-17 (underlined sequence)
Cellular localization: predicted secreted
Summary: TNF-inducible member of hyaluronate binding protein family, related to
CD44--

Please replace the paragraph beginning at page 144, line 9, with the following:

--ACK5 Protein sequence (SEQ ID NO:79):
Gene name: Von Willebrand factor; Coagulation factor VIII
Unigene number: Hs.110802
Probeset Accession #: M10321
Protein Accession #: NP\_000543
Signal peptide: predicted 1-22 (underlined sequence)
Cellular localization: predicted secreted--

Please replace the paragraph beginning at page 144, line 66, with the following:

--AAC7 protein sequence (SEQ ID NO:80):

Gene name: KIAA1294 protein

Probeset Accession #: AA432248

Protein Accession #: BAA92532

Cellular localization: predicted nuclear protein

PFAM prediction: 22-153 Band 41 domain (underlined seq). A number of cytoskeletal-associated proteins that associate with various proteins at the interface between the plasma membrane and the cytoskeleton contain a conserved N-terminal domain of about 150 amino-acid residues.--

Please replace the paragraph beginning at page 145, line 28, with the following:

--ACG8 Protein sequence (SEQ ID NO:81):
Gené name: ubiquitin E3 ligase SMURF2
Unigene number: Hs.21806 (3'UTR only)
Probeset Accession #: AA398243
Protein Accession #: AF301463\_1
Cellular Localization: predicted cytoplasmic
Summary: Smurf2 Is a Ubiquitin E3 Ligase Mediating Proteasome-dependent
Degradation of Smad2 in Transforming Growth Factor-beta Signaling--

MURRAY *et al.* Application No.: 09/784,356

Page 24

Please replace the paragraph beginning at page 145, line 52, with the following:

--ACH1 Protein sequence (SEQ ID NO:82):

Gene name: EST

Unigene number: Hs.30089

Probeset Accession #: AA410480 CAT cluster#: cluster 96816\_1

Summary: predicted open reading frame--

Please replace the paragraph beginning at page 145, line 65, with the following:

--ACJ2 Protein sequence (SEQ ID NO:83):

Gene name: Complement component C1q receptor

Unigene number: Hs.97199

Probeset Accession #: AA487558 Protein Accession #: NP 036204

Signal sequence: 1-17 (first underlined sequence)

Transmemrane domain: 589-605 (second underlined sequence)

Cellular localization: This gene encodes a predicted type I membrane protein. Summary: This protein acts as a receptor for complement protein C1q, mannose-

binding lectin, and pulmonary surfactant protein A. This protein is a

functional receptor involved in ligand-mediated enhancement of phagocytosis.--

Please replace the paragraph beginning at page 146, line 22, with the following:

--ACJ3 Protein sequence (SEQ ID NO:84):

Gene name: FLT1/vascular endothelial growth factor receptor

Unigene number: Hs.138671

Probeset Accession #: AA047437

Transmembrane domain: predicted 764-780 (underlined sequence) Cellular Localization: predicted cell surface tyrosine kinase-

Please replace the paragraph beginning at page 146, line 54, with the following:

--ACJ9 Protein sequence (SEQ ID NO:85):

Gene name: Purine nucleoside phosphorylase

Unigene number: Hs.75514 Probeset Accession #: K02574 Protein Accession #: CAA25320

Cellular Localization: predicted cytoplasmic

Summary: likely to catalyze the reversible phosphorolytic cleavage of purine

ribonucleosides and 2'-deoxyribonucleosides--

MURRAY *et al.* Application No.: 09/784,356

Page 25

Please replace the paragraph beginning at page 147, line 2, with the following:

--ACK4 Protein sequence (SEQ ID NO:86):

Gene name: EST

Probeset Accession #: R68763

Predicted amino acid seq: FGENESH exon prediction on BAC clone AC009414

Predicted nuclear target motifs: from 25 (4) RRRP (SEQ ID NO:87) (underlined);

176 (5) RRRR (SEQ ID NO:88) (underlined); 177 (5) RRRR (SEQ ID NO:88)

(underlined; 239 (5) KRKK (SEQ ID NO:89) (underlined); 399 (4) PPRARRT (SEQ ID NO:90) (underlined); 400 (5) PRARRTE (SEQ ID NO:91) (underlined)

Cellular localization: predicted nuclear--

Please replace the paragraph beginning at page 147, line 22, with the following:

--AAA8 Protein sequence (SEQ ID NO:92):

Gene name: ETL protein, with extended open reading frame

Unigene number: Hs.57958
Probeset Accession #: D58024
Protein Accession #: AAG33021

Transmembrane domains: predicted 454-470, 486-502, 511-527, 528-544, 556-572,

600-616, 642-661, 672-689 (underlined sequences)

Extended sequence: Residues 1-564 were added to the sequence in AAG33021

Cellular Localization: predicted cell surface serpentine receptor--

Please replace the paragraph beginning at page 147, line 46, with the following:

-- AAC6 Protein sequence (SEQ ID NO:93):

Gene name: EST

Unigene number: Hs.134797
Probeset Accession #: AA025351
Protein accession #: BAB14599

Signal sequence: predicted 1-24 (first underlined sequence)

extended sequence: second underlined sequence-

Please replace the paragraph beginning at page 148, line 4, with the following:

-- ACH7 Protein sequence (SEQ ID NO:94):

Gene name: EST

Unigene number: Hs.3807

Probeset Accession #: AA292694

BAC Accession #: AL161751

FGENESH predicted aa seq: 1-647; based on BAC clone AL161751--

Application No.: 09/784,356

Page 26

Please replace the paragraph beginning at page 148, line 24, with the following:

--AAD4 Protein sequence (SEQ ID NO:95):

Gene name: ERG

Unigene number: Hs.45514

Probeset Accession #: R32894

Protein Accession #: AAA52398

Signal sequence: none

Transmembrane domains: none

PFAM domains: predicted Ets-domain 294-373; SAM\_PNT: 122-206 Summary: ERG2 is a sequence-specific DNA-binding protein.--

Please replace the paragraph beginning at page 148, line 44, with the following:

-- AAD5 Protein sequence (SEQ ID NO:96):

Gene name: activin A receptor type II-like 1 (ALK-1)

Unigene number: Hs.172670 Probeset Accession #: T57112 Protein Accession #: NP\_000011 Signal sequence: predicted 1-21

Transmembrane domain: predicted 119-135 PFAM domains: predicted pkinase 204-489

Summary: Type Ia membrane protein; receptor tyrosine kinase-

Please replace the paragraph beginning at page 149, line 21, with the following:

--ACA4 Protein sequence (SEQ ID NO:97):

Gene name: alpha satellite junction DNA sequence

Unigene number: Hs.247946 Probeset Accession #: M21305 Protein Accession #: AAA88020

Signal sequence: none

Transmembrane domains: none

PFAM domains: none--

Application No.: 09/784,356

Page 27

Please replace the paragraph beginning at page 149, line 33, with the following:

--ACG6 Protein sequence (SEQ ID NO:98):

Gene name: intercellular adhesion molecule 2 (ICAM2)

Unigene number: Hs.83733 Probeset Accession #: M32334 Protein Accession #: NP 000864 Signal sequence: predicted 1-21

Transmembrane domain: predicted 224-248

PFAM domains: predicted 41-98, 127-197; immunoglobulin-like C2-type domains Summary: a predicted Type Ia membrane protein; it plays a role in cell adhesion

and is the ligand for the LFA-1 protein. ICAM2 is also called CD102.--

Please replace the paragraph beginning at page 149, line 51, with the following:

--ACG7 Protein sequence (SEQ ID NO:99):

Gene name: Cadherin 5, VE-cadherin (CDH5)

Unigene number: Hs.76206 Probeset Accession #: X79981 Protein Accession #: NP\_001786 Signal sequence: predicted 1-27

Transmembrane domain: predicted 604-620

PFAM domains: Cadherin domains predicted 53-141, 156-249, 263-364, 377-470, and

487-576

Summary: Likely a Type I membrane protein. Cadherins are calcium-dependent adhesive proteins that mediate cell-to-cell interaction. VE-cadherin is associated with intercellular junctions .--

Please replace the paragraph beginning at page 150, line 12, with the following:

--ACG9 Protein sequence (SEQ ID NO:100): Gene name: lysyl oxidase-like 2 (LOXL2)

Unigene number: Hs.83354 Probeset Accession #: U89942 Protein Accession #: NP 002309 Signal sequence: predicted 1-25

Transmembrane domains: none predicted

PFAM domains: scavenger receptor cysteine-rich domains predicted 68-159, 203-

238, 336-425, 439-528; Lysyl oxidase predicted 548-749.

Summary: Likely a secreted protein. Lysyl oxidase is a copper-dependent amine oxidase that belongs to a heterogeneous family of enzymes that oxidize primary amine substrates to reactive aldehydesm, acting on the extracellular matrix substrates, e.g., collagen and elastin.--

Application No.: 09/784,356

Page 28

Please replace the paragraph beginning at page 150, line 41, with the following:

--ACH2 Protein sequence (SEQ ID NO:101):

Gene name: TIE tyrosine-protein kinase
Unigene number: Hs.78824
Probeset Accession #: X60957
Protein Accession #: NP\_005415
Signal sequence: predicted 1-21
Transmembrane domain: predicted 770-786
PFAM domains: laminin-EGF predicted 234-267; FN3 predicted 460-520, 548-632, and 644-729; tyrosine\_kinase predicted 839-1107
Summary: Likely a Type Ia membrane protein; TIE is a tyrosine-kinase receptor with an unknown ligand; its expression is likely necessary for normal blood vessel development.--

Please replace the paragraph beginning at page 151, line 7, with the following:

```
--ACH3 Protein sequence (SEQ ID NO:102):

Gene name: placental growth factor (PGF; PlGF1; VEGF-related protein)

Unigene number: Hs.2894

Probeset Accession #: X54936

Protein Accession #: NP_002623

Signal sequence: predicted 1-21

Transmembrane domain: none predicted

PFAM domains: PDGF predicted 52-130

Summary: Likely a secreted protein; likely regulates angiogenesis by interacting with FLT1 and FLK1.--
```

Please replace the paragraph beginning at page 151, line 23, with the following:

```
--ACH4 Protein sequence (SEQ ID NO:103):

Gene name: nidogen 2 (NID2)

Unigene number: Hs.82733

Probeset Accession #: D86425

Protein Accession #: NP_031387

Signal sequence: predicted 1-30

Transmembrane domain: none predicted

PFAM domains: EGF-like_domains predicted 489-524, 764-800, 806-843, 853-891, and 897-930; thyroglobulin_repeats predicted 941-1006, and 1020-1085;

LDL_receptor_repeats predicted 1155-1197, 1199-1240, and 1242-1285.

Summary: A secreted protein; NID2 likely interacts with collagens I and IV and laminin-1 to promote cell adhesion to the basement membrane.--
```

MURRAY et al. Application No.: 09/784,356

Page 29

Please replace the paragraph beginning at page 151, line 61, with the following:

--ACH5 Protein sequence (SEQ ID NO:104):

Gene name: SNL (singed-like; sea urchin fascin homolog-like)

Unigene number: Hs.118400 Probeset Accession #: U03057 Protein Accession #: NP 003079 Signal sequence: none identified Transmembrane domain: none identified

PFAM domains: none identified

Summary: a cytoplasmic, actin-bundling protein that is likely to be involved in the assembly of actin filament bundles present in microspikes, membrane ruffles,

and stress fibers--

Please replace the paragraph beginning at page 152, line 16, with the following:

--ACH6 Protein sequence (SEQ ID NO:105):

Gene name: endothelial protein C receptor (EPCR; PROCR)

Unigene number: Hs.82353 Probeset Accession #: L35545 Protein Accession #: NP 006395 Signal sequence: predicted 1-17 Transmembrane domain: predicted 211-227

PFAM domains: none identified

Summary: a Type Ia membrane protein, EPCR likely binds to [thrombin]-activated Protein C, a vitamin K-dependent serine protease zymogen necessary for blood

coagulation .--

Please replace the paragraph beginning at page 152, line 34, with the following:

--ACH8 Protein sequence (SEQ ID NO:106):

Gene name: melanoma adhesion molecule (MCAM; MUC18)

Unigene number: Hs.211579 Probeset Accession #: D51069 Protein Accession #: NP 006491 Signal sequence: predicted 1-17

Transmembrane domain: predicted 559-575

PFAM domains: immunoglobulin\_domains predicted 264-324, and 356-410.

Summary: a Type Ia membrane protein, associated with tumor progression and the development of metastasis in human malignant melanoma, and may play a role in neural crest cells during embryonic development .--

Application No.: 09/784,356

Page 30

Please replace the paragraph beginning at page 152, line 59, with the following:

--ACH9 Protein sequence (SEQ ID NO:107):

Gene name: endothelin-1 (EDN1)

Unigene number: Hs.2271
Probeset Accession #: J05008
Protein Accession #: NP\_001946
Signal sequence: predicted 1-17
Transmembrane domain: none predicted

PFAM domains: Endothelin domains predicted 59-73, and 108-129.

Summary: a secreted zymogen; the active protein is likely a 26-amino acid peptide with potent mammalian vasoconstrictor activity; it is necessary for

normal vessel development .--

Please replace the paragraph beginning at page 153, line 11, with the following:

--ACJ1 Protein sequence (SEQ ID NO:108):

Gene name: BMX non-receptor tyrosine kinase

Unigene number: Hs.27372
Probeset Accession #: X83107
Protein Accession #: NP\_001712
Signal sequence: none identified
Transmembrane domain: none identified

PFAM domains: plektrin\_homology\_domain predicted 6-111; SH2 domain predicted

294-383; protein kinase domain predicted 417-663

Summary: a cytoplasmic protein, it likely plays a role in the growth and differentiation of hematopoietic cells; it is known to also be expressed in

endothelial cells .--

Please replace the paragraph beginning at page 153, line 38, with the following:

--ACJ4 Protein sequence (SEQ ID NO:109):

Gene name: prostaglandin G/H synthase 2 (COX-2; PGHS-2)

Unigene number: Hs.196384

Probeset Accession #: D28235

Protein Accession #: NP\_000954

Signal sequence: predicted 1-17

Transmembrane domain: none identified

PFAM domains: EGF-like\_domain predicted 18-55.

Summary: a microsomal enzyme; COX-2 is the therapeutic target of the

nonsteroidal anti-inflammatory drugs (NSAIDs), such as aspirin.--

MURRAY *et al.* Application No.: 09/784,356

Page 31

Please replace the paragraph beginning at page 153, line 62, with the following:

--ACJ6 Protein sequence (SEQ ID NO:110): Gene name: SEC14-like-1

Unigene number: Hs.75232
Probeset Accession #: D67029
Protein Accession #: NP\_002994
Signal sequence: none identified
Transmembrane domain: none identified

PFAM domains: none identified Summary: a cytoplasmic protein--

Please replace the paragraph beginning at page 154, line 18, with the following:

--ACJ8 Protein sequence (SEQ ID NO:111):

Gene name: intercellular adhesion molecule 1 (ICAM1; CD54)

Unigene number: Hs.168383
Probeset Accession #: M24283
Protein Accession #: NP\_000192
Signal sequence: predicted 1-27

Transmembrane domain: predicted 481-497

PFAM domains: immunoglobulin\_domains predicted 128-188, and 325-373. Summary: a Type 1a membrane protein; ICAM1 is typically expressed on endothelial cells and cells of the immune system; ICAM1 binds to integrins of type CD11a/CD18, or CD11b/CD18; ICAM1 is also exploited by Rhinovirus as a receptor.--

Please replace the paragraph beginning at page 154, line 41, with the following:

--ACK3 Protein sequence (SEQ ID NO:112):

Gene name: angiopoietin 1 receptor (TIE-2; TEK)

Unigene number: Hs.89640 Probeset Accession #: L06139 Protein Accession #: NP\_000450 Signal sequence: predicted 1-18

Transmembrane domain: predicted 746-770

PFAM domains: immunoglobulin\_domains predicted 44-102, 370-424; EGF\_like\_domains predicted 210-252, 254-299, and 301-341; FN3\_domains predicted 444-536, 541-634, and 638-732; protein kinase domain predicted 824-1096.

Summary: a Type 1a membrane protein; it is expressed almost exclusively in endothelial cells in mice, rats, and humans; the ligand for this receptor is angiopoietin-1; defects in TEK are associated with inherited venous malformations; the TEK signaling pathway appears to be critical for endothelial cell-smooth muscle cell communication in venous morphogenesis.--

Application No.: 09/784,356

Page 32

Please replace the paragraph beginning at page 155, line 10, with the following:

--PZA6 Protein sequence (SEQ ID NO:113):

Gene name: prostate differentiation factor (PLAB; MIC-1)

Unigene number: Hs.116577 Probeset Accession #: AB000584 Protein Accession #: NP\_004855 Signal sequence: predicted 1-29

Transmembrane domain: none identified

PFAM domains: TGF-beta domain predicted 211-308.

Summary: a secreted protein; its exact function is unclear; it inhibits proliferation of primitive hematopoietic progenitors; it inhibits activation of macrophages; it is highly expressed in placenta and in serum of pregnant women; it may promote fetal survival by suppressing the production of maternally-

derived proinflammatory cytokines within the uterus .--

Please replace the paragraph beginning at page 155, line 32, with the following:

--AAD2 Protein sequence (SEQ ID NO:114):

Gene name: Thrombospondin-1 Unigene number: Hs.87409 Probeset Accession #: AA232645 Protein Accession #: NP 003237.1

Signal sequence: predicted 1-18 (first underlined sequence)

Transmembrane Domain: none identified

Summary: Thrombospondin is a large modular glycoprotein component of the extracellular matrix and contains a variety of distinct domains, including three repeating subunits (types I, II, and III) that share homology to an assortment of other proteins.--

Please replace the paragraph beginning at page 155, line 66, with the following:

--AAD9 protein sequence (SEQ ID NO:115):

Gene name: LIM homeobox protein cofactor (CLIM-1)

Unigene number: Hs.4980 Probeset Accession #: F13782 Protein Accession #: AAC83552

Pfam: LIM bind

Transmembrane Domain: none identifed

Summary: The LIM homeodomain (LIM-HD) proteins, which contain two tandem LIM domains followed by a homeodomain, are critical transcriptional regulators of embryonic development. The LIM domain is a conserved cysteine-rich zinc-binding motif found in LIM-HD proteins, cytoskeletal components, LIM kinases, and other proteins. LIM domains are protein-protein interaction motifs, can inhibit binding of LIM-HD proteins to DNA, and can negatively regulate LIM-HD protein function .--

Application No.: 09/784,356

Page 33

Please replace the paragraph beginning at page 156, line 21, with the following:

--AAE1 protein sequence (SEQ ID NO:116):

Gene name: guanine nucleotide binding protein 11

Unigene number: Hs.83381
Probeset Accession #: U31384
Protein Accession #: NP 004117.1

Pfam: G-gamma; CAAX motif (farnesylation site) prediction underlined Summary: The G gamma proteins are a component of the trimeric G-proteins that interact with cell surface receptors. The G protein beta and gamma subunits directly regulate the activities of various enzymes and ion channels after receptor ligation. Unlike most of the other known gamma subunits, gamma 11 is modified by a farnesyl group and is not capable of interacting with beta 2.--

Please replace the paragraph beginning at page 156, line 37, with the following:

--AAE2 protein sequence (SEQ ID NO:117):

Gene name: Transcription factor 4 (Immunoglobulin transcription factor 2)(ITF-

2) (SL3-3 Enhancer factor 2) (SEF-2)

Unigene number: Hs.289068
Probeset Accession #: M74719
Protein Accession #: NP 003190.1

Pfam: HLH domain prediction underlined

Summary: Transcription factor 4 is a helix-loop-helix (HLH) protein which belongs to a family of nuclear proteins, designated SL3-3 enhancer factors 2 (SEF2), that interact with an Ephrussi box-like motif within the glucocorticoid response element in the enhancer of the murine leukemia virus SL3-3. Various cell types display differences both in the sets of SEF2-DNA complexes formed and in their amounts. Molecular analysis of cDNA clones show the existence of multiple related mRNA species containing alternative coding regions, which are most probably a result of differential splicing.—

Please replace the paragraph beginning at page 156, line 68, with the following:

--AAE4 protein sequence (SEQ ID NO:118):

Gene name: phosphatidylcholine 2-acylhydrolase

Unigene number: Hs.211587
Probeset Accession #: M68874
Protein Accession #: AAA60105.1

Pfam: PLA2 B, C2 domain prediction underlined

Summary: Phospholipases A2 (PLA2s) play a key role in inflammatory processes through production of precursors of eicosanoids and platelet-activating factor. PLA2 is a 100 kd protein that contains a structural element homologous to the C2 region of protein kinase C.--

Application No.: 09/784,356

Page 34

Please replace the paragraph beginning at page 157, line 26, with the following:

--ACA1 protein sequence (SEQ ID NO:119):

Gene name: tissue factor pathway inhibitor 2 TFPI2, placental protein 5 (PP5)

Unigene number: Hs.78045 Probeset Accession #: D29992 Protein Accession #: BAA06272.1

Pfam: Kunitz BPTI

Signal sequence: underlined

Summary: ACA1 is a serine proteinase inhibitor that was originally purified from conditioned medium of the human glioblastoma cell line T98G. ACA1 is identical to placental protein 5 (PP5) and TFPI2, a placenta-derived glycoprotein with serine proteinase inhibitor activity. PP5 belongs to the Kunitz-type serine proteinase inhibitor family, having three putative Kunitztype inhibitor domains .--

Please replace the paragraph beginning at page 157, line 45, with the following:

--ACB8 protein sequence (SEQ ID NO:120):

Gene name: myosin X

Unigene number: Hs.61638 Probeset Accession #: N77151

Protein Accession #: NP\_036466

Pfam: myosin head, IQ (calmodulin binding motif), PH, MyTH4

Summary: Myosins are molecular motors that move along filamentous actin. Seven classes of myosin are expressed in vertebrates: conventional myosin, or myosin-II, as well as the 6 unconventional myosin classes-I, -V, -VI, -VII, -IX, and -

X . --

Please replace the paragraph beginning at page 158, line 24, with the following:

--ACC3 protein sequence (SEQ ID NO:121):

Gene name: calcitonin receptor-like (CALCRL)

Unigene number: Hs.152175 Probeset Accession #: L76380 Protein Accession #: NP 005786.1

Pfam: 7TM 2 (7 transmembrane receptor (Secretin family))

Transmembrane domains: predictions underlined

Signal sequence: first underlined region

Summary: Calcitonin gene-related peptide (CGRP) is a neuropeptide with diverse biological effects including potent vasodilator activity. The human CGRP1 receptor shares significant peptide sequence homology with the human calcitonin receptor, a member of the G-protein-coupled receptor superfamily. expression in 293 (HEK 293) cells produces specific, high affinity binding sites for CGRP. Exposure of these cells to CGRP results in a 60-fold increase in cAMP production . --

hak

MURRAY et al.

PATENT

Application No.: 09/784,356

Page 35

Please replace the paragraph beginning at page 158, line 49, with the following:

-- ACC5 protein sequence (SEQ ID NO:122):

Gene name: Selectin E (endothelial adhesion molecule 1)

Unigene number: Hs.89546
Probeset Accession #: M24736
Protein Accession #: NP\_000441.1

Pfam: lectin c, EGF like domain, sushi (SCR domain)

Signal sequence: first underlined region

Transmembrane domain: second underlined region

Summary: Focal adhesion of leukocytes to the blood vessel lining is a key step in inflammation and certain vascular disease processes. Endothelial leukocyte adhesion molecule-1 (ELAM-1), a cell surface glycoprotein expressed by cytokine-activated endothelium, mediates the adhesion of blood neutrophils. The primary sequence of ELAM-1 predicts an amino-terminal lectin-like domain, an EGF domain, and six tandem repetitive motifs (about 60 amino acids each) related to those found in complement regulatory proteins. A similar domain structure is also found in the MEL-14 lymphocyte cell surface homing receptor, and in granule-membrane protein 140, a membrane glycoprotein of platelet and endothelial secretory granules that can be rapidly mobilized (less than 5 minutes) to the cell surface by thrombin and other stimuli. Thus, ELAM-1 may be a member of a nascent gene family of cell surface molecules involved in the regulation of inflammatory and immunological events at the interface of vessel wall and blood.--

Please replace the paragraph beginning at page 159, line 17, with the following:

--ACC8 protein sequence (SEQ ID NO:123):

Gene name: Chemokine (C-X-C motif), receptor 4 (fusin)

Unigene number: Hs.89414
Probeset Accession #: L06797
Protein Accession #: NP\_003458.1

Pfam: 7TM 1 (7 transmembrane receptor (rhodopsin family))

Signal sequence: none identified

Transmembrane domains: predictions underlined

Summary: The chemokine receptor CXCR4 (also designated fusin and LESTR) is a

cofactor for fusion and entry of T cell-tropic strains of HIV-1.--

MURRAY et al. Application No.: 09/784,356

Page 36

Please replace the paragraph beginning at page 159, line 36, with the following:

--ACF2 protein sequence (SEQ ID NO:124):

Gene name: Endothelial cell-specific molecule 1

Unigene number: Hs.41716
Probeset Accession #: X89426
Protein Accession #: NP\_008967.1
Signal sequence: underlined

Pfam: IGFBP (Insulin-like growth factor binding proteins)

Summary: Human endothelial cell-specific molecule (called ESM-1) was cloned from a human umbilical vein endothelial cell (HUVEC) cDNA library. Constitutive ESM-1 gene expression is seen in HUVECs but not in the other human cell lines. The cDNA sequence contains an open reading frame of 552 nucleotides and a 1398-nucleotide 3'-untranslated region including several domains involved in mRNA instability and five putative polyadenylation consensus sequences. The deduced 184-amino acid sequence defines a cysteine-rich protein with a functional NH2-terminal hydrophobic signal sequence.--

Please replace the paragraph beginning at page 159, line 58, with the following:

--ACF4 protein sequence (SEQ ID NO:125):

Gene name: P53-responsive gene 2 similar to D.melanogaster peroxidasin(U11052)

Unigene number: Hs.118893 Probeset Accession #: D86983 Protein Accession #: BAA13219

Pfam: LRRNT (Leucine rich repeat N-terminal domain), LRR (Leucine Rich Repeat),

LRRCT (Leucine rich repeat C-terminal domain), Ig (immunoglobulin domain),

Peroxidase, VWC (von Willebrand factor type C domain)

Summary: ACF4 is a gene originally identified from KG-1 cell and brain cDNA

libraries.--

Application No.: 09/784,356

Page 37

Please replace the paragraph beginning at page 160, line 28, with the following:

--ACF5 protein sequence (SEQ ID NO:126):

Gene name: Mitogen-activated protein kinase kinase kinase 4

Unigene number: Hs.3628

Probeset Accession #: N54067 Protein Accession #: NP\_004825.1

Pfam: pkinase (Eukaryotic protein kinase domain), CNH domain Summary: The yeast serine/threonine kinase STE20 activates a signaling cascade that includes STE11 (mitogen-activated protein kinase kinase kinase), STE7 (mitogen-activated protein kinase kinase), and FUS3/KSS1 (mitogen-activated protein kinase) in response to signals from both Cdc42 and the heterotrimeric G proteins associated with transmembrane pheromone receptors. ACF5 is a human cDNA encoding a protein kinase homologous to STE20. This protein kinase, also designated HPK/GCK-like kinase (HGK), has nucleotide sequences that encode an open reading frame of 1165 amino acids with 11 kinase subdomains. HGK is a serine/threonine protein kinase that specifically activated the c-Jun N-terminal kinase (JNK) signaling pathway when transfected into 293T cells, but does not stimulate either the extracellular signal-regulated kinase or p38 kinase pathway. HGK also increased AP-1-mediated transcriptional activity in vivo. HGK may be a novel activator of the JNK pathway. The cascade may look like this: HGK -> TAK1 -> MKK4, MKK7 -> JNK kinase cascade, which may mediate the TNFalpha signaling pathway .--

Please replace the paragraph beginning at page 161, line 4, with the following:

--ACF8 protein sequence (SEQ ID NO:127):

Gene name: Phospholipase A2, group IVC (cytosolic, calcium-independent)

Unigene number: Hs.18858

Probeset Accession #: AA054087 Protein Accession #: NP\_003697.1

Pfam: none identified

Summary: ACF8 is a membrane-bound, calcium-independent PLA2, named cPLA2-gamma. The sequence encodes a 541-amino acid protein containing a domain with significant homology to the catalytic domain of the 85-kDa cPLA2 (cPLA2-alpha). cPLA2-gamma does not contain the regulatory calcium-dependent lipid binding (CaLB) domain found in cPLA2-alpha. cPLA2-gamma does contain two consensus motifs for lipid modification, a prenylation motif (-CCLA) at the C terminus and a myristoylation site at the N terminus. cPLA2-gamma demonstrates a preference for arachidonic acid at the sn-2 position of phosphatidylcholine as compared with palmitic acid. cPLA2-gamma encodes a 3-kilobase message, which is highly expressed in heart and skeletal muscle, suggesting a specific role in these tissues.--

MURRAY et al. Application No.: 09/784,356

Page 38

Please replace the paragraph beginning at page 161, line 34, with the following:

--ACG1 protein sequence (SEQ ID NO:128):

Gene name: Carbohydrate (chondroitin 6/keratan) sulfotransferase 1

Unigene number: Hs.104576 Probeset Accession #: AA868063 Protein Accession #: NP\_003645.1

Pfam: none identified

Summary: Chondroitin 6-sulfotransferase (C6ST) is the key enzyme in the biosynthesis of chondroitin 6-sulfate, a glycosaminoglycan implicated in chondrogenesis, neoplasia, atherosclerosis, and other processes. C6ST catalyzes the transfer of sulfate from 3'-phosphoadenosine 5'-phosphosulfate to carbon 6 of the N-acetylgalactosamine residues of chondroitin.--

Please replace the paragraph beginning at page 161, line 55, with the following:

--ACG5 protein sequence (SEQ ID NO:129):

Gene name: Multimerin Unigene number: Hs.268107 Probeset Accession #: U27109 Protein Accession #: AAC52065

Signal sequence: prediction underlined

Pfam: EGF-like domain, Clq domain

Summary: Multimerin is a massive, soluble protein found in platelets and in the endothelium of blood vessels. Multimerin is composed of varying sized, disulfide-linked multimers, the smallest of which is a homotrimer. Multimerin is a factor V/Va-binding protein and may function as a carrier protein for platelet factor V. Northern analyses show a 4.7-kilobase transcript in cultured endothelial cells, a megakaryocytic cell line, platelets, and highly vascular tissues. The multimerin cDNA can encode a protein of 1228 amino acids with the probable signal peptide cleavage site between amino acids 19 and 20. The protein is predicted to be hydrophilic and to contain 23 N-glycosylation sites. The adhesive motif RGDS (Arg-Gly-Asp-Ser) and an epidermal growth factor-like domain were identified. Multimerin contains a probable coiled-coil structures in the central portion of its sequence. Additionally, the carboxyl-terminal region of multimerin resembles the globular, non-collagen-like, carboxyl-terminal domains of several other trimeric proteins, including complement Clq and collagens type VIII and X.--

Please replace the paragraph beginning at page 162, line 32, with the following:

--ACC6 protein sequence (SEQ ID NO:131):

Gene name: Homo sapiens cDNA FLJ11502 fis, clone HEMBA1002102, weakly similar

to ANKRYIN

Unigene number: Hs.213194
Probeset Accession #: AA187101
Protein Accession #: none
Pfam: ankyrin repeats--

MURRAY et al.

Application No.: 09/784,356

Page 39

Please replace the paragraph beginning at page 162, line 45, with the following:

--ACC7 protein sequence (SEQ ID NO:132):

Gene name: Human RAL A gene Uniquene number: Hs.6906

Probeset Accession #: AA083572 cluster

Protein Accession #: P11233

Pfam: ras

Features: CAAX motif is underlined

Summary: The RALA gene encodes a low molecular mass ras-like GTP-binding protein that shares about 50% similarity with the ras proteins. GTP-binding proteins mediate the transmembrane signaling initiated by the occupancy of certain cell surface receptors. The RALA gene maps to 7p22-p15.--

Please replace the paragraph beginning at page 162, line 63, with the following:

--ACC9 protein sequence (SEQ ID NO:133):

Gene name: KIAA0955 protein Unigene number: Hs.10031

Probeset Accession #: AA027168 Protein Accession #: BAA76799.1

Pfam: CARD (Caspase recruitment domain)

Summary: Gene was originally isolated as a brain cDNA. The coding region contains a CARD domain, suggesting involvement in apoptotic signaling pathways.--

Please replace the paragraph beginning at page 163, line 14, with the following:

--ACF6 Protein sequence (SEQ ID NO:134):

Gene name: Homo sapiens cDNA FLJ10669 fis, clone NT2RP2006275, weakly similar

to Microtubule-associated protein 1B [CONTAINS: LIGHT CHAIN LC1]

Unigene number: Hs.66048

Probeset Accession #: AA609717 Protein Accession #: BAA91743.1

Pfam: none identified

Summary: The cDNA for FLJ10669 was originally isolated from NT2 neuronal precursor cells (teratocarcinoma cell line) after 2-weeks of retinoic acid (RA) treatment. The protein sequence has similarity to microtubule-associated protein 1B (MAP-1B), suggesting a function for ACF6 in the regulating the cytoskeleton.--

Please insert the accompanying paper copy of the Sequence Listing, page numbers 1 to 154, at the end of the application.

Application No.: 09/784,356

Page 40

## **REMARKS**

Applicants request entry of this amendment in adherence with 37 C.F.R. §§1.821 to 1.825. If the Examiner believes a telephone conference would expedite prosecution of this application, please telephone the undersigned at 415-576-0200. eM

Respectfully submitted,

Reg. No. 44,879

TOWNSEND and TOWNSEND and CREW LLP Two Embarcadero Center, 8<sup>th</sup> Floor San Francisco, California 94111-3834

Tel: (415) 576-0200 Fax: (415) 576-0300

JML:dmw

Application No.: 09/784,356

Page 41

## VERSION WITH MARKINGS TO SHOW CHANGES MADE

## In the Specification:

Paragraph beginning at line 23 of page 42 has been amended as follows:

Various tag polypeptides and their respective antibodies are well known in the art. Examples include poly-histidine (poly-his) or poly-histidine-glycine (poly-his-gly) tags; HIS6 (SEQ ID NO:135) and metal chelation tags, the flu HA tag polypeptide and its antibody 12CA5 [Field et al., Mol. Cell. Biol., 8:2159-2165 (1988)]; the c-myc tag and the 8F9, 3C7, 6E10, G4, B7 and 9E10 antibodies thereto [Evan et al., Molecular and Cellular Biology, 5:3610-3616 (1985)]; and the Herpes Simplex virus glycoprotein D (gD) tag and its antibody [Paborsky et al., Protein Engineering, 3(6):547-553 (1990)]. Other tag polypeptides include the Flag-peptide [Hopp et al., BioTechnology, 6:1204-1210 (1988)]; the KT3 epitope peptide [Martin et al., Science, 255:192-194 (1992)]; tubulin epitope peptide [Skinner et al., J. Biol. Chem., 266:15163-15166 (1991)]; and the T7 gene 10 protein peptide tag [Lutz-Freyermuth et al., Proc. Natl. Acad. Sci. USA, 87:6393-6397 (1990)].

Paragraph beginning at line 3 of page 82 has been amended as follows:

AAA4 DNA sequence (SEQ ID NO:1):

Gene name: CGI-100 protein Unigene number: Hs.275253

Probeset Accession #: AA089688

Nucleic Acid Accession #: NM 016040 cluster

Coding sequence: 142-831 (predicted start/stop codons underlined)

Paragraph beginning at line 32 of page 82 has been amended as follows:

AAA7 DNA sequence (SEQ ID NO:2):

Gene name: Endothelial differentiation, sphingolipid G-protein-coupled

receptor, 1 (EDG1)

Unigene number: Hs.154210 Probeset Accession #: M31210

Nucleic Acid Accession #: NM 001400 cluster

Coding sequence: 251-1396 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 42

Paragraph beginning at line 20 of page 83 has been amended as follows:

AAB3 DNA sequence (SEQ ID NO:3):

Gene name: Solute carrier family 20 (phosphate transporter), member 1, Human

leukaemia virus receptor 1 (GLVR1)

Unigene number: Hs.78452 Probeset Accession #: L20859

Nucleic Acid Accession #: NM\_005415 cluster

Coding sequence: predicted 371-2410 (predicted start/stop codons underlined)

Paragraph beginning at line 16 of page 84 has been amended as follows:

AAB4 DNA sequence (SEQ ID NO:4):

Gene name: Matrix metalloproteinase 10 (stromelysin 2)

Unigene number: Hs.2258

Probeset Accession #: X07820

Nucleic Acid Accession #: NM 002425

Coding sequence: predicted 23-1453 (predicted start/stop codons underlined)

Paragraph beginning at line 55 of page 84 has been amended as follows:

AAB6 DNA sequence (SEQ ID NO:5):

Gene name: Podocalyxin-like
Unigene number: Hs.16426
Probeset Accession #: U97519

Nucleic Acid Accession #: NM 005397 cluster

Coding sequence: 251-1837 (predicted start/stop codons underlined)

Paragraph beginning at line 26 of page 86 has been amended as follows:

AAB8 DNA sequence (SEQ ID NO:6):

Gene name: EGF-containing fibulin-like extracellular matrix protein 1

Unigene number: Hs.76224 Probeset Accession #: U03877

Nucleic Acid Accession #: NM\_004105 Transcript variant 1

Coding sequence: 150-1631 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 43

Paragraph beginning at line 13 of page 87 has been amended as follows:

AAB9 DNA sequence (SEQ ID NO:7):

Gene name: Melanoma adhesion molecule, MUC 18 glycoprotein

Unigene number: Hs.211579 Probeset Accession #: M28882

Nucleic Acid Accession #: NM 006500 cluster

Coding sequence: 27-1967 (predicted start/stop codons underlined)

Paragraph beginning at line 14 of page 88 has been amended as follows:

AAC1 DNA sequence (SEQ ID NO:8):

Gene name: Matrix metalloproteinase 1 (interstitial collagenase)

Unigene number: Hs.83169 Probeset Accession #: X54925

Nucleic Acid Accession #: NM\_002421 cluster

Coding sequence: 69-1478 (predicted start/stop codons underlined)

Paragraph beginning at line 56 of page 88 has been amended as follows:

AAC3 DNA sequence (SEQ ID NO:9):

Gene name: Branched chain aminotransferase 1, cytosolic

Unigene number: Hs.157205

Probeset Accession #: AA423987

Nucleic Acid Accession #: NM 005504 cluster

Coding sequence: 1-1155 (predicted start/stop codons underlined)

Paragraph beginning at line 17 of page 89 has been amended as follows:

ACG4 DNA sequence: SEQ ID NO:10):

Gene name: Pentaxin-related gene, rapidly induced by IL-1 beta

Unigene number: Hs.2050

Probeset Accession #: M31166

Nucleic Acid Accession #: NM\_002852 cluster

Coding sequence: 68-1213 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 44

Ti. 1.2 TT: m

Hand the

Paragraph beginning at line 57 of page 89 has been amended as follows:

ACK5 DNA sequence (SEQ ID NO:11):

Gene name: Von Willebrand factor; Coagulation factor VIII

Unigene number: Hs.110802 Probeset Accession #: M10321

Nucleic Acid Accession #: NM 000552

Coding sequence: 311-8752 (predicted start/stop codons underlined)

**PATENT** 

Paragraph beginning at line 11 of page 92 has been amended as follows:

AAC7 DNA sequence (SEQ ID NO:12):

Gene name: KIAA1294 protein Probeset Accession #: AA432248

Nucleic Acid Accession #: AB037715

Coding sequence: 370-3489 (predicted start/stop codons underlined)

Paragraph beginning at line 65 of page 93 has been amended as follows:

ACG8 DNA sequence (SEQ ID NO:13):

Gene name: ubiquitin E3 ligase SMURF2 Unigene number: Hs.21806 (3'UTR only)

Probeset Accession #: AA398243

Nucleic Acid Accession #: AF301463 cluster

Coding sequence: 9-2255 (predicted start/stop codons underlined)

Paragraph beginning at line 44 of page 94 has been amended as follows:

ACH1 DNA sequence (SEQ ID NO:14):

Gene name: EST

Unigene number: Hs.30089

Probeset Accession #: AA410480

CAT cluster#: 96816 1

Coding sequence: Partial sequence, possible frameshift. Predicted stop codon

underlined.

Page 45

Paragraph beginning at line 2 of page 95 has been amended as follows:

ACJ2 DNA sequence (SEQ ID NO:15):

Gene name: Complement component Clq receptor

Unigene number: Hs.97199

Probeset Accession #: AA487558

Nucleic Acid Accession #: NM 012072

Coding sequence: 149-2107. Predicted start/stop codons underlined

Paragraph beginning at line 55 of page 96 has been amended as follows:

ACJ3 DNA sequence (SEQ ID NO:16):

Gene name: FLT1/vascular endothelial growth factor receptor

Unigene number: Hs.138671 Probeset Accession #: AA047437

Nucleic Acid Accession #: NM 002019

Coding sequence: 250-4266 (predicted start/stop codons underlined)

Paragraph beginning at line 56 of page 98 has been amended as follows:

ACJ9 DNA sequence (SEQ ID NO:17):

Gene name: Purine nucleoside phosphorylase

Unigene number: Hs.75514 Probeset Accession #: K02574

Nucleic acid Accession #: X00737 cluster

Coding sequence: 110-979 (predicted start/stop codons underlined)

Paragraph beginning at line 21 of page 99 has been amended as follows:

ACK4 DNA sequence (SEQ ID NOS:18 and 19):

Gene name: EST

Uniquene number: Hs.265499 Probeset Accession #: R68763 CAT cluster#: Cluster 46668 2

Sequence: Both the EST corresponding to the probeset accession and exon prediction; number and the CAT cluster align with the Homo sapiens BAC clone AC009414 RP11-490M8. Using FGENESH, 2 exons predicted on this BAC clone

upstream of the probeset.

Predicted exon 1: bases 5808-5837 of BAC clone AC009414

Paragraph beginning at line 58 of page 99 has been amended as follows:

Predicted ACK4 gene seq (predicted start/stop codons underlined) (SEQ ID NO:19):

Application No.: 09/784,356 Page 46

Paragraph beginning at line 20 of page 100 has been amended as follows:

AAA8 DNA sequence (SEQ ID NO:20):

Gene name: ETL protein, with extended open reading frame

Unigene number: Hs.57958
Probeset Accession #: D58024
Nucleotide Accession #: AF192403

Coding sequence: 151-2136. Underlined sequences correspond to extended

sequence not included in AF192403.

Paragraph beginning at line 11 of page 101 has been amended as follows:

AAC6 DNA sequence (SEQ ID NO:21):

Gene name: Homo sapiens cDNA FLJ13465 fis, clone PLACE1003493, weakly similar

to endothelial cell multimerin precursor

Unigene number: Hs.134797
Probeset Accession #: AA025351
Nucleotide Accession #: AK023527
Coding sequence: predicted 75-2921

Extended sequence: 729-3465 (underlined sequence)

Paragraph beginning at line 18 of page 102 has been amended as follows:

ACH7 DNA sequence (SEQ ID NOS:22-24):

Gene name: ESTs

Ţ

The state of the s

Unigene number: Hs.3807

Probeset Accession #: AA292694

BAC Accession #: AL161751

FGENESH predicted exons: FGENESH predicts 2 exons on the minus strand of

AL161751 upstream of the ACH7 probeset.

Paragraph beginning at line 26 of page 102 has been amended as follows:

FGENESH predicted exon 1 (SEQ ID NO:22):

Paragraph beginning at line 32 of page 102 has been amended as follows:

FGENESH predicted exon 2 (SEQ ID NO:23):

Application No.: 09/784,356

Page 47

Paragraph beginning at line 63 of page 102 has been amended as follows:

ACH7 predicted coding seq (predicted start/stop codons underlined)  $\underline{\text{(SEQ ID NO:24)}}$ :

Paragraph beginning at line 31 of page 103 has been amended as follows:

AAD3 DNA sequence (SEQ ID NO:25):

Gene name: ESTs

Unigene number: Hs.17404
Probeset Accession #: N39584
Nucleic Acid Accession #: N39584

Coding sequence: no identified ORF; possible frameshifts

Paragraph beginning at line 64 of page 103 has been amended as follows:

AAD4 DNA sequence (SEQ ID NO:26):

Gene name: ERG

Unigene number: Hs.279477 / Hs.45514

Probeset Accession #: R32894 Nucleic Acid Accession #: M17254

Coding sequence: 257-1645 (predicted start/stop codons underlined)

Paragraph beginning at line 58 of page 104 has been amended as follows:

AAD5 DNA sequence (SEQ ID NO:27):

Gene name: activin A receptor type II-like 1 (ALK-1)

Unigene number: Hs.8881 / Hs.172670

Probeset Accession #: T57112

Nucleic Acid Accession #: NM\_000020

Coding sequence: 283-1794 (predicted start/stop codons underlined)

Paragraph beginning at line 3 of page 106 has been amended as follows:

AAD8 DNA sequence (SEQ ID NO:28):

Gene name: ESTs

Unigene number: Hs.144953

Probeset Accession #: AA404418 Nucleic Acid Accession #: n/a

Coding sequence: no ORF identified; possible frameshifts

Application No.: 09/784,356

Page 48

Paragraph beginning at line 39 of page 106 has been amended as follows:

ACA2 DNA sequence (SEQ ID NO:29):

Gene name: EST

Unigene number: Hs.16450

Probeset Accession #: AA478778

Nucleic Acid Accession #: AA478778

Coding sequence: no ORF identified; possible frameshifts

Paragraph beginning at line 1 of page 107 has been amended as follows:

ACA4 DNA sequence (SEQ ID NO:30):

Gene name: alpha satellite junction DNA sequence

Unigene number: Hs.247946
Probeset Accession #: M21305
Nucleic Acid Accession #: M21305

Coding sequence: 1-165 (predicted start/stop codons underlined)

Paragraph beginning at line 13 of page 107 has been amended as follows:

ACG6 DNA sequence (SEQ ID NO:31):

Gene name: intercellular adhesion molecule 2 (ICAM2)

Unigene number: Hs.83733
Probeset Accession #: M32334

Nucleic Acid Accession #: NM 000873

Coding sequence: 63-890 (predicted start/stop codons underlined)

Paragraph beginning at line 40 of page 107 has been amended as follows:

ACG7 DNA sequence (SEQ ID NO:32):

Gene name: Cadherin 5, VE-cadherin (CDH5)

Unigene number: Hs.76206 Probeset Accession #: X79981

Nucleic Acid Accession #: NM 001795

Coding sequence: 25-2379 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 49

Paragraph beginning at line 48 of page 108 has been amended as follows:

ACG9 DNA sequence (SEQ ID NO:33):

Gene name: lysyl oxidase-like 2 (LOXL2)

Unigene number: Hs.83354 Probeset Accession #: U89942

Nucleic Acid Accession #: NM\_002318 cluster

Coding sequence: 248-2572 (predicted start/stop codons underlined)

PATENT

Paragraph beginning at line 47 of page 109 has been amended as follows:

ACH2 DNA sequence (SEQ ID NO:34):

Gene name: TIE tyrosine-protein kinase

Unigene number: Hs.78824 Probeset Accession #: X60957

Nucleic Acid Accession #: NM 005424 cluster

Coding sequence: 37-3452 (predicted start/stop codons underlined)

Paragraph beginning at line 53 of page 110 has been amended as follows:

ACH3 DNA sequence (SEQ ID NO:35):

Gene name: placental growth factor (PGF; PlGF1; VEGF-related protein)

Unigene number: Hs.2894

Probeset Accession #: X54936

Nucleic Acid Accession #: NM\_002632 cluster

Coding sequence: 322-768 (predicted start/stop codons underlined)

Paragraph beginning at line 22 of page 111 has been amended as follows:

ACH4 DNA sequence (SEQ ID NO:36):

Gene name: nidogen 2 (NID2) Unigene number: Hs.82733 Probeset Accession #: D86425

Nucleic Acid Accession #: NM 007361 cluster

Coding sequence: 1-4131 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 50

Paragraph beginning at line 44 of page 112 has been amended as follows:

ACH5 DNA sequence (SEQ ID NO:37):

Gene name: SNL (singed-like; sea urchin fascin homolog-like)

Unigene number: Hs.118400 Probeset Accession #: U03057

Nucleic Acid Accession #: NM 003088

Coding sequence: 112-1593 (predicted start/stop codons underlined)

Paragraph beginning at line 32 of page 113 has been amended as follows:

ACH6 DNA sequence (SEQ ID NO:38):

Gene name: endothelial protein C receptor (EPCR; PROCR)

Unigene number: Hs.82353 Probeset Accession #: L35545

Nucleic Acid Accession #: NM 006404

Coding sequence: 25-741 (predicted start/stop codons underlined)

Paragraph beginning at line 63 of page 113 has been amended as follows:

ACH8 DNA sequence (SEQ ID NO:39):

Gene name: melanoma adhesion molecule (MCAM; MUC18)

Unigene number: Hs.211579 Probeset Accession #: D51069

Nucleic Acid Accession #: NM 006500

Coding sequence: 27-1967 (predicted start and stop codons underlined)

Paragraph beginning at line 64 of page 114 has been amended as follows:

ACH9 DNA sequence (SEQ ID NO:40):

Gene name: endothelin-1 (EDN1)

Unigene number: Hs.2271

Probeset Accession #: J05008

Nucleic Acid Accession #: NM 001955

Coding sequence: 337-975 (predicted start/stop codons underlined)

II. 5 

Application No.: 09/784,356

Page 51

Paragraph beginning at line 26 of page 115 has been amended as follows:

ACJ1 DNA sequence (SEQ ID NO:41):

Gene name: BMX non-receptor tyrosine kinase

Unigene number: Hs.27372 Probeset Accession #: X83107

Nucleic Acid Accession #: NM 001721

Coding sequence: 34-2061 (predicted start/stop codons underlined)

Paragraph beginning at line 8 of page 116 has been amended as follows:

ACJ4 DNA sequence (SEQ ID NO:42):

Gene name: prostaglandin G/H synthase 2 (COX-2; PGHS-2)

Unigene number: Hs.196384 Probeset Accession #: D28235

Nucleic Acid Accession #: NM 000963

Coding sequence: 135-1949 (predicted start/stop codons underlined)

Paragraph beginning at line 24 of page 117 has been amended as follows:

ACJ6 DNA sequence (SEQ ID NO:43):

Gene name: SEC14-like-1
Unigene number: Hs.75232
Probeset Accession #: D67029

Nucleic Acid Accession #: NM 003003

Coding sequence: 304-2451 (predicted start/stop codons underlined)

Paragraph beginning at line 56 of page 118 has been amended as follows:

ACJ8 DNA sequence (SEQ ID NO:44):

Gene name: intercellular adhesion molecule 1 (ICAM1; CD54)

Unigene number: Hs.168383 Probeset Accession #: M24283

Nucleic Acid Accession #: NM 000201

Coding sequence: 58-1656 (predicted start/stop codons underlined)

Application No.: 09/784,356

Page 52

Paragraph beginning at line 47 of page 119 has been amended as follows:

ACK3 DNA sequence (SEQ ID NO:45):

Gene name: angiopoietin 1 receptor (TIE-2; TEK)

Unigene number: Hs.89640 Probeset Accession #: L06139

Nucleic Acid Accession #: NM 000459

Coding sequence: 149-3523 (predicted start/stop codons underlined)

Paragraph beginning at line 57 of page 120 has been amended as follows:

PZA6 DNA sequence (SEQ ID NO:46):

Gene name: prostate differentiation factor (PLAB; MIC-1)

Unigene number: Hs.116577

Probeset Accession #: AB000584 Nucleic Acid Accession #: NM 004864

Coding sequence: 26-952 (predicted start/stop codons underlined)

Paragraph beginning at line 19 of page 121 has been amended as follows:

AAC8 DNA sequence (SEQ ID NO:47):

Gene name: none

Uniquene number: Hs.6682

Probeset Accession #: AA227926 Nucleic Acid Accession #: none

Coding sequence: no ORF identified, possible frameshifts

Paragraph beginning at line 12 of page 122 has been amended as follows:

AAD2 DNA sequence (SEQ ID NO:48):

Gene name: Thrombospondin-1 Unigene number: Hs.87409

Probeset Accession #: AA232645 Nucleic Acid Accession #: NM 003246

Coding sequence: 112-3624 (predicted start/stop codons underlined)

111

Application No.: 09/784,356

Page 53

Paragraph beginning at line 49 of page 123 has been amended as follows:

AAD9 DNA sequence (SEQ ID NO:49):

Gene name: LIM homeobox protein cofactor (CLIM-1)

Unigene number: Hs.4980 Probeset Accession #: F13782

Nucleic Acid Accession #: AF047337

Coding sequence: 110-1231(predicted start/stop codons underlined)

Paragraph beginning at line 29 of page 124 has been amended as follows:

AAE1 DNA sequence (SEQ ID NO:50):

Gene name: guanine nucleotide binding protein 11

Unigene number: Hs.83381 Probeset Accession #: U31384

Nucleic Acid Accession #: NM\_004126.1

Coding sequence: 108-329 (predicted start/stop codons underlined)

Paragraph beginning at line 49 of page 124 has been amended as follows:

AAE2 DNA sequence (SEQ ID NO:51):

Gene name: Transcription factor 4 (immunoglobulin transcription factor 2) (ITF-

2) (SL3-3 Enhancer factor 2) (SEF-2)

Unigene number: Hs.289068
Probeset Accession #: M74719

Nucleic Acid Accession #: NM 003199.1

Coding sequence: 200-2203 (predicted start/stop codons underlined)

Paragraph beginning at line 33 of page 125 has been amended as follows:

AAE4 DNA sequence (SEQ ID NO:52):

Gene name: phosphatidylcholine 2-acylhydrolase

Unigene number: Hs.211587
Probeset Accession #: M68874
Nucleic Acid Accession #: M68874

Coding sequence: 139-2388 (predicted start/stop codons underlined)

Page 54

Paragraph beginning at line 22 of page 126 has been amended as follows:

ACA1 DNA sequence (SEQ ID NO:53):

Gene name: tissue factor pathway inhibitor 2 TFPI2, placental protein 5 (PP5)

Unigene number: Hs.78045 Probeset Accession #: D29992

Nucleic Acid Accession #: D29992.1

Coding sequence: 57-764 (predicted start/stop codons underlined)

Paragraph beginning at line 51 of page 126 has been amended as follows:

ACB8 DNA sequence (SEQ ID NO:54):

Gene name: myosin X

Unigene number: Hs.61638

Probeset Accession #: N77151

Nucleic Acid Accession #: NM 012334

Coding sequence: 223-6399 (predicted start/stop codons underlined)

Paragraph beginning at line 54 of page 128 has been amended as follows:

ACC3 DNA sequence (SEQ ID NO:55):

Gene name: calcitonin receptor-like (CALCRL)

Unigene number: Hs.152175 Probeset Accession #: L76380

Nucleic Acid Accession #: NM 005795

Coding sequence: 555-1940 (predicted start/stop codons underlined)

Paragraph beginning at line 46 of page 129 has been amended as follows:

ACC4 DNA sequence (SEQ ID NO:56):

Gene name: Homo sapiens mRNA; cDNA DKFZp586E1624

Unigene number: Hs.94030

Probeset Accession #: AA452000

Nucleic Acid Accession #: AL110152.1

Coding sequence: no ORF identified, possible frameshifts

With this E But High 

Application No.: 09/784,356

Page 55

Paragraph beginning at line 10 of page 130 has been amended as follows:

ACC5 DNA sequence (SEQ ID NO:57):

Gene name: Selectin E (endothelial adhesion molecule 1)

Unigene number: Hs.89546 Probeset Accession #: M24736

Nucleic Acid Accession #: NM\_000450

Coding sequence: 117-1949 (predicted start/stop codons underlined)

Paragraph beginning at line 15 of page 131 has been amended as follows:

ACC8 DNA sequence (SEQ ID NO:58):

Gene name: Chemokine (C-X-C motif), receptor 4 (fusin)

Unigene number: Hs.89414

Probeset Accession #: L06797

Nucleic Acid Accession #: NM 003467

Coding sequence: 89-1147 (predicted start/stop codons underlined)

Paragraph beginning at line 52 of page 131 has been amended as follows:

ACF2 DNA sequence (SEQ ID NO:59):

Gene name: Endothelial cell-specific molecule 1

Unigene number: Hs.41716 Probeset Accession #: X89426

Nucleic Acid Accession #: NM 007036

Coding sequence: 56-610 (predicted start/stop codons underlined)

Paragraph beginning at line 27 of page 132 has been amended as follows:

ACF4 DNA sequence (SEQ ID NO:60):

Gene name: P53-responsive gene 2 similar to D.melanogaster peroxidasin(U11052)

Unigene number: Hs.118893 Probeset Accession #: D86983 Nucleic Acid Accession #: D86983

Coding sequence: 1-4491 (predicted stop codon underlined, sequence is open at

5' end)

Application No.: 09/784,356

Page 56

Paragraph beginning at line 61 of page 133 has been amended as follows:

ACF5 DNA sequence (SEQ ID NO:61):

Gene name: Mitogen-activated protein kinase kinase kinase 4

Unigene number: Hs.3628

Probeset Accession #: N54067

Nucleic Acid Accession #: NM 004834

Coding sequence: 80-3577 (predicted start/stop codons underlined)

Paragraph beginning at line 67 of page 134 has been amended as follows:

ACF8 DNA sequence (SEQ ID NO:62):

Gene name: Phospholipase A2, group IVC (cytosolic, calcium-independent)

Unigene number: Hs.18858

Probeset Accession #: AA054087

Nucleic Acid Accession #: NM\_003706

Coding sequence: 310-1935 (predicted start/stop codons underlined)

Paragraph beginning at line 50 of page 135 has been amended as follows:

ACG1 DNA sequence (SEQ ID NO:63):

Gene name: Carbohydrate (chondroitin 6/keratan) sulfotransferase 1

Unigene number: Hs.104576

Probeset Accession #: AA868063

Nucleic Acid Accession #: NM 003654

Coding sequence: 367-1602 (predicted start/stop codons underlined)

Paragraph beginning at line 32 of page 136 has been amended as follows:

ACG5 DNA sequence (SEQ ID NO:64):

Gene name: Multimerin

Unigene number: Hs.268107

Probeset Accession #: U27109

Nucleic Acid Accession #: U27109.1

Coding sequence: 72-3758 (predicted start/stop codons underlined)

Paragraph beginning at line 43 of page 137 has been amended as follows:

ACC6 DNA sequence (SEQ ID NO:65):

Gene name: Homo sapiens cDNA FLJ11502 fis, clone HEMBA1002102, weakly similar

to ANKRYIN

Unigene number: Hs.213194 Probeset Accession #: AA187101 Nucleic Acid Accession #: AK021564

Coding sequence: 1-450 (predicted stop codon underlined, 5'end sequence is

open)

Paragraph beginning at line 13 of page 138 has been amended as follows:

ACC7 DNA sequence (SEQ ID NO:66):

Gene name: Human RAL A gene

Unigene number: Hs.6906

Probeset Accession #: AA083572

Nucleic Acid Accession #: contig of X15014.1 and AK026850

Coding sequence: 1-621 (predicted start/stop codons underlined)

Paragraph beginning at line 65 of page 138 has been amended as follows:

ACC9 DNA sequence (SEQ ID NO:67):

Gene name: KIAA0955 protein Uniquene number: Hs.10031

Probeset Accession #: AA027168 Nucleic Acid Accession #: AB023172

Coding sequence: 314-1609 (predicted start/stop codons underlined)

Paragraph beginning at line 23 of page 140 has been amended as follows:

ACF6 DNA sequence (SEQ ID NO:68):

Gene name: Homo sapiens cDNA FLJ10669 fis, clone NT2RP2006275, weakly similar

to Microtubule-associated protein 1B [CONTAINS: LIGHT CHAIN LC1]

Unigene number: Hs.66048

Probeset Accession #: AA609717 Nucleic Acid Accession #: AK001531

Coding sequence: 176-2194 (predicted start/stop codons underlined)

MURRAY et al. Application No.: 09/784,356

Page 58

Paragraph beginning at line 4 of page 141 has been amended as follows:

AAA4 Protein sequence+ (SEQ ID NO:69):

Gene name: CGI-100 protein

Unigene number: Hs.275253

Probeset Accession #: AA089688

Protein Accession #: NP 057124

Signal sequence: predicted 1-23 (first underlined sequence)

Transmembrane Domain: predicted 201-217 (second underlined sequence)

emp24/gp25L/p24 domain: predicted 13-227

Summary: gp25L/emp24/p24 protein family members of the cis-Golgi network bind both COP I and II coatomer. Members of this family are implicated in bringing cargo forward from the ER and binding to coat proteins by their cytoplasmic domains.

Paragraph beginning at line 22 of page 141 has been amended as follows:

AAA7 Protein sequence: (SEQ ID NO:70):

Gene name: Endothelial differentiation, sphingolipid G-protein-coupled

receptor, 1 (EDG1)

Unigene number: Hs.154210 Probeset Accession #: M31210 Protein Accession#: NP\_001391

7 Transmembrane Domains: predicted 50-71, 92-110, 122-140, 160-177, 201-222,

251-269, 281-301 (underlined sequences)

Summary: Endothelial differentiation, sphingolipid G-protein-coupled receptor,

1 may regulate the differentiation of endothelial cells. It binds the

sphingolipid metabolite, sphingosine-1-phosphate, which may function as a second

messenger in cell proliferation and survival.

Paragraph beginning at line 44 of page 141 has been amended as follows:

AAB3 Protein sequence: (SEQ ID NO:71)

Gene name: Solute carrier family 20 (phosphate transporter), member 1, Human

leukaemia virus receptor 1 (GLVR1)

Unigene number: Hs.78452 Probeset Accession #: L20859 Protein Accession #: NP\_005406

Transmembrane domains: predicted 24-40, 62-78, 164-180, 198-214, 232-248, 513-

529, 562-578, 604-620, 655-671

Cellular Localization: Likely a Type IIIa membrane protein (Ncyt Cexo)

Application No.: 09/784,356

Page 59

Paragraph beginning at line 68 of page 141 has been amended as follows:

AAB4 Protein sequence: (SEQ ID NO:72):

Gene name: Matrix metalloproteinase 10 (stromelysin 2)

Unigene number: Hs.2258
Probeset Accession #: X07820
Protein Accession #: NP\_002416

Signal sequence: predicted 1-17 (underlined sequence)

Cellular Localization: predicted secreted

Paragraph beginning at line 18 of page 142 has been amended as follows:

AAB6 Protein sequence (SEQ ID NO:73):

Gene name: Podocalyxin-like Unigene number: Hs.16426 Probeset Accession #: U97519 Protein Accession #: NP 005388

Transmembrane domain: predicted 432-448 (underlined sequence)
Cellular Localization: predicted Type Ia membrane protein (Nexo)

Paragraph beginning at line 37 of page 142 has been amended as follows:

AAB8 Protein sequence: (SEQ ID NO:74):

Gene name: EGF-containing fibulin-like extracellular matrix protein 1

Unigene number: Hs.76224 Probeset Accession #: U03877

Protein Accession #: NP 004096 Variant 1

Signal sequence: predicted 1-17 (underlined sequence)

Summary: This gene spans approximately 18 kb of genomic DNA and consists of 12 exons. Two transcripts with distinct 5' UTR have been described; the resulting proteins have distinct N-terminal amino acid sequences. Translation initiation from internal methionine residues was observed with in vitro translation. A signal peptide sequence is predicted for translation initiation sites 1, 2, and 4. The protein isoforms contain 5 or 6 calcium-binding EGF2 domains and 5 or 6 EGF2 domains. Mutations in this gene cause the retinal disease Malattia Leventinese. Transcript Variant: This variant (1) has a distinct 5' UTR and N-terminal protein sequence as compared to variant 2.

Application No.: 09/784,356

Page 60

Paragraph beginning at line 64 of page 142 has been amended as follows:

AAB9 Protein sequence+ (SEQ ID NO:75):

Gene name: Melanoma adhesion molecule, MUC 18 glycoprotein

Unigene number: Hs.211579
Probeset Accession #: M28882
Protein Accession #: NP\_006491

Signal sequence: predicted 1-17 (first underlined sequence)

Transmembrane domain: predicted 559-575 (second underlined sequence) Cellular localization: predicted Type Ia membrane protein (Nexo)

Paragraph beginning at line 18 of page 143 has been amended as follows:

AAC1 Protein sequence: (SEQ ID NO:76):

Gene name: Matrix metalloproteinase 1 (interstitial collagenase)

Unigene number: Hs.83169
Probeset Accession #: X54925
Protein Accession #: NP 002412

Signal sequence: predicted 1-19 (underlined sequence) Cellular localization: predicted secreted protein

Paragraph beginning at line 36 of page 143 has been amended as follows:

AAC3 Protein sequence: (SEQ ID NO:77):

Gene name: Branched chain aminotransferase 1, cytosolic

Unigene number: Hs.157205

Probeset Accession #: AA423987
Protein Accession #: NP\_005495
Cellular Localization: cytolasmic

Summary: The lack of the cytosolic enzyme branched-chain amino acid transaminase (BCT) causes cell growth inhibition. There may be at least 2 different clinical disorders due to a defect of branched-chain amino acid transamination: hypervalinemia and hyperleucine-isoleucinemia. Since there are 2 distinct BCATs, mitochondrial and cytosolic, it is possible that one is mutant in each of these 2 conditions.

Application No.: 09/784,356

Page 61

Paragraph beginning at line 58 of page 143 has been amended as follows:

ACG4 Protein sequence: (SEQ ID NO:78):

Gene name: Pentaxin-related gene, rapidly induced by IL-1 beta

Unigene number: Hs.2050 Probeset Accession #: M31166 Protein Accession #: NP 002843

Signal sequence: predicted 1-17 (underlined sequence)

Cellular localization: predicted secreted

Summary: TNF-inducible member of hyaluronate binding protein family, related to

CD44

Paragraph beginning at line 9 of page 144 has been amended as follows:

ACK5 Protein sequence: (SEQ ID NO:79):

Gene name: Von Willebrand factor; Coagulation factor VIII

Unigene number: Hs.110802 Probeset Accession #: M10321 Protein Accession #: NP\_000543

Signal peptide: predicted 1-22 (underlined sequence)

Cellular localization: predicted secreted

Paragraph beginning at line 66 of page 144 has been amended as follows:

AAC7 protein sequence: (SEQ ID NO:80):

Gene name: KIAA1294 protein
Probeset Accession #: AA432248
Protein Accession #: BAA92532

Cellular localization: predicted nuclear protein

PFAM prediction: 22-153 Band 41 domain (underlined seq). A number of cytoskeletal-associated proteins that associate with various proteins at the interface between the plasma membrane and the cytoskeleton contain a conserved N-terminal domain of about 150 amino-acid residues.

Paragraph beginning at line 28 of page 145 has been amended as follows:

ACG8 Protein sequence: (SEQ ID NO:81):

Gene name: ubiquitin E3 ligase SMURF2

Unigene number: Hs.21806 (3'UTR only)

Probeset Accession #: AA398243 Protein Accession #: AF301463\_1

Cellular Localization: predicted cytoplasmic

Summary: Smurf2 Is a Ubiquitin E3 Ligase Mediating Proteasome-dependent

Degradation of Smad2 in Transforming Growth Factor-beta Signaling

Application No.: 09/784,356

Page 62

Paragraph beginning at line 52 of page 145 has been amended as follows:

ACH1 Protein sequence+ (SEQ ID NO:82):

Gene name: EST

Unigene number: Hs.30089

Probeset Accession #: AA410480 CAT cluster#: cluster 96816\_1

Summary: predicted open reading frame

Paragraph beginning at line 65 of page 145 has been amended as follows:

ACJ2 Protein sequence: (SEQ ID NO:83):

Gene name: Complement component Clq receptor

Unigene number: Hs.97199

Probeset Accession #: AA487558 Protein Accession #: NP 036204

Signal sequence: 1-17 (first underlined sequence)

Transmemrane domain: 589-605 (second underlined sequence)

Cellular localization: This gene encodes a predicted type I membrane protein. Summary: This protein acts as a receptor for complement protein C1q, mannose-

binding lectin, and pulmonary surfactant protein A. This protein is a

functional receptor involved in ligand-mediated enhancement of phagocytosis.

Paragraph beginning at line 22 of page 146 has been amended as follows:

ACJ3 Protein sequence+ (SEQ ID NO:84):

Gene name: FLT1/vascular endothelial growth factor receptor

Unigene number: Hs.138671 Probeset Accession #: AA047437

Transmembrane domain: predicted 764-780 (underlined sequence) Cellular Localization: predicted cell surface tyrosine kinase

Paragraph beginning at line 54 of page 146 has been amended as follows:

ACJ9 Protein sequence: (SEQ ID NO:85):

Gene name: Purine nucleoside phosphorylase

Unigene number: Hs.75514
Probeset Accession #: K02574
Protein Accession #: CAA25320

Cellular Localization: predicted cytoplasmic

Summary: likely to catalyze the reversible phosphorolytic cleavage of purine

ribonucleosides and 2'-deoxyribonucleosides

Application No.: 09/784,356

Page 63

Paragraph beginning at line 2 of page 147 has been amended as follows:

ACK4 Protein sequence (SEQ ID NO:86):

Gene name: EST

Probeset Accession #: R68763

Predicted amino acid seq: FGENESH exon prediction on BAC clone AC009414 Predicted nuclear target motifs: from 25 (4) RRRP (SEQ ID NO:87) (underlined); 176 (5) RRRR (SEQ ID NO:88) (underlined); 177 (5) RRRR (SEQ ID NO:88) (underlined; 239 (5) KRKK (SEQ ID NO:89) (underlined); 399 (4) PPRARRT (SEQ ID NO:90) (underlined); 400 (5) PRARRTE (SEQ ID NO:91) (underlined) Cellular localization: predicted nuclear

Paragraph beginning at line 22 of page 147 has been amended as follows:

AAA8 Protein sequence (SEQ ID NO:92):

Gene name: ETL protein, with extended open reading frame

Unigene number: Hs.57958 Probeset Accession #: D58024 Protein Accession #: AAG33021

Transmembrane domains: predicted 454-470, 486-502, 511-527, 528-544, 556-572,

600-616, 642-661, 672-689 (underlined sequences)

Extended sequence: Residues 1-564 were added to the sequence in AAG33021

Cellular Localization: predicted cell surface serpentine receptor

Paragraph beginning at line 46 of page 147 has been amended as follows:

AAC6 Protein sequence: (SEQ ID NO:93):

Gene name: EST

Uniquene number: Hs.134797 Probeset Accession #: AA025351 Protein accession #: BAB14599

Signal sequence: predicted 1-24 (first underlined sequence)

extended sequence: second underlined sequence

Paragraph beginning at line 4 of page 148 has been amended as follows:

ACH7 Protein sequence: (SEQ ID NO:94):

Gene name: EST

Uniquene number: Hs.3807

Probeset Accession #: AA292694

BAC Accession #: AL161751

FGENESH predicted aa seq: 1-647; based on BAC clone AL161751

Application No.: 09/784,356

Page 64

Paragraph beginning at line 24 of page 148 has been amended as follows:

AAD4 Protein sequence (SEQ ID NO:95):

Gene name: ERG

Unigene number: Hs.45514
Probeset Accession #: R32894
Protein Accession #: AAA52398

Signal sequence: none

Transmembrane domains: none

PFAM domains: predicted Ets-domain 294-373; SAM\_PNT: 122-206 Summary: ERG2 is a sequence-specific DNA-binding protein.

Paragraph beginning at line 44 of page 148 has been amended as follows:

AAD5 Protein sequence (SEQ ID NO:96):

Gene name: activin A receptor type II-like 1 (ALK-1)

Unigene number: Hs.172670
Probeset Accession #: T57112
Protein Accession #: NP\_000011
Signal sequence: predicted 1-21

Transmembrane domain: predicted 119-135 PFAM domains: predicted pkinase 204-489

Summary: Type Ia membrane protein; receptor tyrosine kinase

Paragraph beginning at line 21 of page 149 has been amended as follows:

ACA4 Protein sequence (SEQ ID NO:97):

Gene name: alpha satellite junction DNA sequence

Unigene number: Hs.247946 Probeset Accession #: M21305 Protein Accession #: AAA88020

Signal sequence: none

Transmembrane domains: none

PFAM domains: none

Application No.: 09/784,356

Page 65

Paragraph beginning at line 33 of page 149 has been amended as follows:

ACG6 Protein sequence (SEQ ID NO:98):

Gene name: intercellular adhesion molecule 2 (ICAM2)

Unigene number: Hs.83733
Probeset Accession #: M32334
Protein Accession #: NP\_000864
Signal sequence: predicted 1-21

Transmembrane domain: predicted 224-248

PFAM domains: predicted 41-98, 127-197; immunoglobulin-like C2-type domains Summary: a predicted Type Ia membrane protein; it plays a role in cell adhesion

and is the ligand for the LFA-1 protein. ICAM2 is also called CD102.

Paragraph beginning at line 51 of page 149 has been amended as follows:

ACG7 Protein sequence (SEQ ID NO:99):

Gene name: Cadherin 5, VE-cadherin (CDH5)

Unigene number: Hs.76206
Probeset Accession #: X79981
Protein Accession #: NP\_001786
Signal sequence: predicted 1-27

Transmembrane domain: predicted 604-620

PFAM domains: Cadherin domains predicted 53-141, 156-249, 263-364, 377-470, and

487-576

Summary: Likely a Type I membrane protein. Cadherins are calcium-dependent adhesive proteins that mediate cell-to-cell interaction. VE-cadherin is associated with intercellular junctions.

Paragraph beginning at line 12 of page 150 has been amended as follows:

ACG9 Protein sequence (SEQ ID NO:100):

Gene name: lysyl oxidase-like 2 (LOXL2)

Unigene number: Hs.83354
Probeset Accession #: U89942
Protein Accession #: NP\_002309
Signal sequence: predicted 1-25

Transmembrane domains: none predicted

PFAM domains: scavenger receptor cysteine-rich domains predicted 68-159, 203-

238, 336-425, 439-528; Lysyl oxidase predicted 548-749.

Summary: Likely a secreted protein. Lysyl oxidase is a copper-dependent amine oxidase that belongs to a heterogeneous family of enzymes that oxidize primary amine substrates to reactive aldehydesm, acting on the extracellular matrix substrates, e.g., collagen and elastin.

Application No.: 09/784,356

Page 66

Paragraph beginning at line 41 of page 150 has been amended as follows:

ACH2 Protein sequence (SEQ ID NO:101): Gene name: TIE tyrosine-protein kinase

Unigene number: Hs.78824 Probeset Accession #: X60957 Protein Accession #: NP 005415 Signal sequence: predicted 1-21

Transmembrane domain: predicted 770-786

PFAM domains: laminin-EGF predicted 234-267; FN3 predicted 460-520, 548-632,

and 644-729; tyrosine kinase predicted 839-1107

Summary: Likely a Type Ia membrane protein; TIE is a tyrosine-kinase receptor with an unknown ligand; its expression is likely necessary for normal blood

vessel development.

Paragraph beginning at line 7 of page 151 has been amended as follows:

ACH3 Protein sequence (SEQ ID NO:102):

Gene name: placental growth factor (PGF; PlGF1; VEGF-related protein)

Unigene number: Hs.2894 Probeset Accession #: X54936 Protein Accession #: NP 002623 Signal sequence: predicted 1-21 Transmembrane domain: none predicted PFAM domains: PDGF predicted 52-130

Summary: Likely a secreted protein; likely regulates angiogenesis by

interacting with FLT1 and FLK1.

Paragraph beginning at line 23 of page 151 has been amended as follows:

ACH4 Protein sequence (SEQ ID NO:103):

Gene name: nidogen 2 (NID2) Unigene number: Hs.82733 Probeset Accession #: D86425 Protein Accession #: NP 031387 Signal sequence: predicted 1-30 Transmembrane domain: none predicted

PFAM domains: EGF-like domains predicted 489-524, 764-800, 806-843, 853-891,

and 897-930; thyroglobulin repeats predicted 941-1006, and 1020-1085; LDL receptor\_repeats predicted 1155-1197, 1199-1240, and 1242-1285.

Summary: A secreted protein; NID2 likely interacts with collagens I and IV and

laminin-1 to promote cell adhesion to the basement membrane.

Application No.: 09/784,356

Page 67

Paragraph beginning at line 61 of page 151 has been amended as follows:

ACH5 Protein sequence (SEQ ID NO:104):

Gene name: SNL (singed-like; sea urchin fascin homolog-like)

Unigene number: Hs.118400
Probeset Accession #: U03057
Protein Accession #: NP\_003079
Signal sequence: none identified
Transmembrane domain: none identified

PFAM domains: none identified

Summary: a cytoplasmic, actin-bundling protein that is likely to be involved in the assembly of actin filament bundles present in microspikes, membrane ruffles,

and stress fibers

Paragraph beginning at line 16 of page 152 has been amended as follows:

ACH6 Protein sequence (SEQ ID NO:105):

Gene name: endothelial protein C receptor (EPCR; PROCR)

Unigene number: Hs.82353
Probeset Accession #: L35545
Protein Accession #: NP\_006395
Signal sequence: predicted 1-17

Transmembrane domain: predicted 211-227

PFAM domains: none identified

Summary: a Type Ia membrane protein, EPCR likely binds to [thrombin]-activated Protein C, a vitamin K-dependent serine protease zymogen necessary for blood coagulation.

Paragraph beginning at line 34 of page 152 has been amended as follows:

ACH8 Protein sequence (SEQ ID NO:106):

Gene name: melanoma adhesion molecule (MCAM; MUC18)

Unigene number: Hs.211579
Probeset Accession #: D51069
Protein Accession #: NP\_006491
Signal sequence: predicted 1-17

Transmembrane domain: predicted 559-575

PFAM domains: immunoglobulin\_domains predicted 264-324, and 356-410.

Summary: a Type Ia membrane protein, associated with tumor progression and the development of metastasis in human malignant melanoma, and may play a role in neural crest cells during embryonic development.

Application No.: 09/784,356

Page 68

Paragraph beginning at line 59 of page 152 has been amended as follows:

ACH9 Protein sequence (SEQ ID NO:107):

Gene name: endothelin-1 (EDN1)

Unigene number: Hs.2271

Probeset Accession #: J05008
Protein Accession #: NP 001946

Signal sequence: predicted 1-17

Transmembrane domain: none predicted

PFAM domains: Endothelin domains predicted 59-73, and 108-129.

Summary: a secreted zymogen; the active protein is likely a 26-amino acid peptide with potent mammalian vasoconstrictor activity; it is necessary for

normal vessel development.

Paragraph beginning at line 11 of page 153 has been amended as follows:

ACJ1 Protein sequence (SEQ ID NO:108):

Gene name: BMX non-receptor tyrosine kinase

Unigene number: Hs.27372

Probeset Accession #: X83107

Protein Accession #: NP\_001712

Signal sequence: none identified

Transmembrane domain: none identified

PFAM domains: plektrin\_homology\_domain predicted 6-111; SH2\_domain predicted

294-383; protein\_kinase\_domain predicted 417-663

Summary: a cytoplasmic protein, it likely plays a role in the growth and differentiation of hematopoietic cells; it is known to also be expressed in

endothelial cells.

Paragraph beginning at line 38 of page 153 has been amended as follows:

ACJ4 Protein sequence (SEQ ID NO:109):

Gene name: prostaglandin G/H synthase 2 (COX-2; PGHS-2)

Unigene number: Hs.196384
Probeset Accession #: D28235
Protein Accession #: NP\_000954
Signal sequence: predicted 1-17

Transmembrane domain: none identified

PFAM domains: EGF-like\_domain predicted 18-55.

Summary: a microsomal enzyme; COX-2 is the therapeutic target of the

nonsteroidal anti-inflammatory drugs (NSAIDs), such as aspirin.

MURRAY et al. Application No.: 09/784,356

Page 69

Paragraph beginning at line 62 of page 153 has been amended as follows:

ACJ6 Protein sequence (SEQ ID NO:110):

Gene name: SEC14-like-1
Unigene number: Hs.75232
Probeset Accession #: D67029
Protein Accession #: NP\_002994
Signal sequence: none identified
Transmembrane domain: none identified
PFAM domains: none identified
Summary: a cytoplasmic protein

Paragraph beginning at line 18 of page 154 has been amended as follows:

ACJ8 Protein sequence (SEQ ID NO:111):

Gene name: intercellular adhesion molecule 1 (ICAM1; CD54)

Unigene number: Hs.168383

Probeset Accession #: M24283

Protein Accession #: NP\_000192

Signal sequence: predicted 1-27

Transmembrane domain: predicted 481-497

PFAM domains: immunoglobulin\_domains predicted 128-188, and 325-373. Summary: a Type la membrane protein; ICAM1 is typically expressed on endothelial cells and cells of the immune system; ICAM1 binds to integrins of type CD11a/CD18, or CD11b/CD18; ICAM1 is also exploited by Rhinovirus as a receptor.

Paragraph beginning at line 41 of page 154 has been amended as follows:

ACK3 Protein sequence (SEQ ID NO:112):

Gene name: angiopoietin 1 receptor (TIE-2; TEK)

Unigene number: Hs.89640
Probeset Accession #: L06139
Protein Accession #: NP\_000450
Signal sequence: predicted 1-18
Transmembrane domain: predicted

Transmembrane domain: predicted 746-770

PFAM domains: immunoglobulin\_domains predicted 44-102, 370-424; EGF\_like\_domains predicted 210-252, 254-299, and 301-341; FN3\_domains predicted 444-536, 541-634, and 638-732; protein\_kinase\_domain predicted 824-1096.

Summary: a Type la membrane protein; it is expressed almost exclusively in endothelial cells in mice, rats, and humans; the ligand for this receptor is angiopoietin-1; defects in TEK are associated with inherited venous malformations; the TEK signaling pathway appears to be critical for endothelial cell-smooth muscle cell communication in venous morphogenesis.

Application No.: 09/784,356

Page 70

Paragraph beginning at line 10 of page 155 has been amended as follows:

PZA6 Protein sequence (SEQ ID NO:113):

Gene name: prostate differentiation factor (PLAB; MIC-1)

Unigene number: Hs.116577
Probeset Accession #: AB000584
Protein Accession #: NP\_004855
Signal sequence: predicted 1-29

Transmembrane domain: none identified

PFAM domains: TGF-beta domain predicted 211-308.

Summary: a secreted protein; its exact function is unclear; it inhibits proliferation of primitive hematopoietic progenitors; it inhibits activation of macrophages; it is highly expressed in placenta and in serum of pregnant women; it may promote fetal survival by suppressing the production of maternally-derived proinflammatory cytokines within the uterus.

Paragraph beginning at line 32 of page 155 has been amended as follows:

AAD2 Protein sequence+ (SEQ ID NO:114):

Gene name: Thrombospondin-1 Unigene number: Hs.87409

Probeset Accession #: AA232645 Protein Accession #: NP\_003237.1

Signal sequence: predicted 1-18 (first underlined sequence)

Transmembrane Domain: none identified

Summary: Thrombospondin is a large modular glycoprotein component of the extracellular matrix and contains a variety of distinct domains, including three repeating subunits (types I, II, and III) that share homology to an assortment of other proteins.

Paragraph beginning at line 66 of page 155 has been amended as follows:

AAD9 protein sequence (SEQ ID NO:115):

Gene name: LIM homeobox protein cofactor (CLIM-1)

Unigene number: Hs.4980

Probeset Accession #: F13782 Protein Accession #: AAC83552

Pfam: LIM bind

Transmembrane Domain: none identifed

Summary: The LIM homeodomain (LIM-HD) proteins, which contain two tandem LIM domains followed by a homeodomain, are critical transcriptional regulators of embryonic development. The LIM domain is a conserved cysteine-rich zinc-binding motif found in LIM-HD proteins, cytoskeletal components, LIM kinases, and other proteins. LIM domains are protein-protein interaction motifs, can inhibit binding of LIM-HD proteins to DNA, and can negatively regulate LIM-HD protein function.

MURRAY et al. Application No.: 09/784,356

Page 71

Paragraph beginning at line 21 of page 156 has been amended as follows:

AAE1 protein sequence (SEQ ID NO:116):

Gene name: guanine nucleotide binding protein 11

Unigene number: Hs.83381
Probeset Accession #: U31384
Protein Accession #: NP\_004117.1

Pfam: G-gamma; CAAX motif (farnesylation site) prediction underlined Summary: The G gamma proteins are a component of the trimeric G-proteins that interact with cell surface receptors. The G protein beta and gamma subunits directly regulate the activities of various enzymes and ion channels after receptor ligation. Unlike most of the other known gamma subunits, gamma 11 is modified by a farnesyl group and is not capable of interacting with beta 2.

Paragraph beginning at line 37 of page 156 has been amended as follows:

AAE2 protein sequence (SEQ ID NO:117):

Gene name: Transcription factor 4 (Immunoglobulin transcription factor 2) (ITF-

2) (SL3-3 Enhancer factor 2) (SEF-2)

Unigene number: Hs.289068
Probeset Accession #: M74719
Protein Accession #: NP 003190.1

Pfam: HLH domain prediction underlined

Summary: Transcription factor 4 is a helix-loop-helix (HLH) protein which belongs to a family of nuclear proteins, designated SL3-3 enhancer factors 2 (SEF2), that interact with an Ephrussi box-like motif within the glucocorticoid response element in the enhancer of the murine leukemia virus SL3-3. Various cell types display differences both in the sets of SEF2-DNA complexes formed and in their amounts. Molecular analysis of cDNA clones show the existence of multiple related mRNA species containing alternative coding regions, which are most probably a result of differential splicing.

Paragraph beginning at line 68 of page 156 has been amended as follows:

AAE4 protein sequence (SEQ ID NO:118):

Gene name: phosphatidylcholine 2-acylhydrolase

Unigene number: Hs.211587
Probeset Accession #: M68874
Protein Accession #: AAA60105.1

Pfam: PLA2 B, C2 domain prediction underlined

Summary: Phospholipases A2 (PLA2s) play a key role in inflammatory processes through production of precursors of eicosanoids and platelet-activating factor. PLA2 is a 100 kd protein that contains a structural element homologous to the C2 region of protein kinase C.

MURRAY et al. Application No.: 09/784,356

Page 72

Paragraph beginning at line 26 of page 157 has been amended as follows:

ACA1 protein sequence (SEQ ID NO:119):

Gene name: tissue factor pathway inhibitor 2 TFPI2, placental protein 5 (PP5)

Unigene number: Hs.78045 Probeset Accession #: D29992 Protein Accession #: BAA06272.1

Pfam: Kunitz BPTI

Signal sequence: underlined

Summary: ACA1 is a serine proteinase inhibitor that was originally purified from conditioned medium of the human glioblastoma cell line T98G. ACA1 is identical to placental protein 5 (PP5) and TFPI2, a placenta-derived glycoprotein with serine proteinase inhibitor activity. PP5 belongs to the Kunitz-type serine proteinase inhibitor family, having three putative Kunitz-type inhibitor domains.

Paragraph beginning at line 45 of page 157 has been amended as follows:

ACB8 protein sequence (SEQ ID NO:120):

Gene name: mvosin X

Unigene number: Hs.61638
Probeset Accession #: N77151
Protein Accession #: NP 036466

Pfam: myosin head, IQ (calmodulin binding motif), PH, MyTH4

Summary: Myosins are molecular motors that move along filamentous actin. Seven classes of myosin are expressed in vertebrates: conventional myosin, or myosin-II, as well as the 6 unconventional myosin classes-I, -V, -VI, -VII, -IX, and -X.

Paragraph beginning at line 24 of page 158 has been amended as follows:

ACC3 protein sequence (SEQ ID NO:121):

Gene name: calcitonin receptor-like (CALCRL)

Unigene number: Hs.152175 Probeset Accession #: L76380 Protein Accession #: NP 005786.1

Pfam: 7TM 2 (7 transmembrane receptor (Secretin family))

Transmembrane domains: predictions underlined

Signal sequence: first underlined region

Summary: Calcitonin gene-related peptide (CGRP) is a neuropeptide with diverse biological effects including potent vasodilator activity. The human CGRP1 receptor shares significant peptide sequence homology with the human calcitonin receptor, a member of the G-protein-coupled receptor superfamily. Stable expression in 293 (HEK 293) cells produces specific, high affinity binding sites for CGRP. Exposure of these cells to CGRP results in a 60-fold increase in cAMP production.

Application No.: 09/784,356

Page 73

Paragraph beginning at line 49 of page 158 has been amended as follows:

ACC5 protein sequence (SEQ ID NO:122):

Gene name: Selectin E (endothelial adhesion molecule 1)

Unigene number: Hs.89546
Probeset Accession #: M24736
Protein Accession #: NP\_000441.1

Pfam: lectin c, EGF like domain, sushi (SCR domain)

Signal sequence: first underlined region

Transmembrane domain: second underlined region

Summary: Focal adhesion of leukocytes to the blood vessel lining is a key step in inflammation and certain vascular disease processes. Endothelial leukocyte adhesion molecule-1 (ELAM-1), a cell surface glycoprotein expressed by cytokine-activated endothelium, mediates the adhesion of blood neutrophils. The primary sequence of ELAM-1 predicts an amino-terminal lectin-like domain, an EGF domain, and six tandem repetitive motifs (about 60 amino acids each) related to those found in complement regulatory proteins. A similar domain structure is also found in the MEL-14 lymphocyte cell surface homing receptor, and in granule-membrane protein 140, a membrane glycoprotein of platelet and endothelial secretory granules that can be rapidly mobilized (less than 5 minutes) to the cell surface by thrombin and other stimuli. Thus, ELAM-1 may be a member of a nascent gene family of cell surface molecules involved in the regulation of inflammatory and immunological events at the interface of vessel wall and blood.

Paragraph beginning at line 17 of page 159 has been amended as follows:

ACC8 protein sequence (SEQ ID NO:123):

Gene name: Chemokine (C-X-C motif), receptor 4 (fusin)

Unigene number: Hs.89414
Probeset Accession #: L06797
Protein Accession #: NP\_003458.1

Pfam: 7TM 1 (7 transmembrane receptor (rhodopsin family))

Signal sequence: none identified

Transmembrane domains: predictions underlined

Summary: The chemokine receptor CXCR4 (also designated fusin and LESTR) is a

cofactor for fusion and entry of T cell-tropic strains of HIV-1.

Application No.: 09/784,356

Page 74

Paragraph beginning at line 36 of page 159 has been amended as follows:

ACF2 protein sequence (SEQ ID NO:124):

Gene name: Endothelial cell-specific molecule 1

Unigene number: Hs.41716
Probeset Accession #: X89426
Protein Accession #: NP\_008967.1
Signal sequence: underlined

Pfam: IGFBP (Insulin-like growth factor binding proteins)

Summary: Human endothelial cell-specific molecule (called ESM-1) was cloned from a human umbilical vein endothelial cell (HUVEC) cDNA library. Constitutive ESM-1 gene expression is seen in HUVECs but not in the other human cell lines. The cDNA sequence contains an open reading frame of 552 nucleotides and a 1398-nucleotide 3'-untranslated region including several domains involved in mRNA instability and five putative polyadenylation consensus sequences. The deduced 184-amino acid sequence defines a cysteine-rich protein with a functional NH2-terminal hydrophobic signal sequence.

Paragraph beginning at line 58 of page 159 has been amended as follows:

ACF4 protein sequence (SEQ ID NO:125):

Gene name: P53-responsive gene 2 similar to D.melanogaster peroxidasin(U11052)

Unigene number: Hs.118893 Probeset Accession #: D86983 Protein Accession #: BAA13219

Pfam: LRRNT (Leucine rich repeat N-terminal domain), LRR (Leucine Rich Repeat),

LRRCT (Leucine rich repeat C-terminal domain), Ig (immunoglobulin domain),

Peroxidase, VWC (von Willebrand factor type C domain)

Summary: ACF4 is a gene originally identified from KG-1 cell and brain cDNA

libraries.

MURRAY et al. Application No.: 09/784,356

Page 75

Paragraph beginning at line 28 of page 160 has been amended as follows:

ACF5 protein sequence (SEQ ID NO:126):

Gene name: Mitogen-activated protein kinase kinase kinase kinase 4

Unigene number: Hs.3628 Probeset Accession #: N54067

Protein Accession #: NP 004825.1

Pfam: pkinase (Eukaryotic protein kinase domain), CNH domain Summary: The yeast serine/threonine kinase STE20 activates a signaling cascade that includes STE11 (mitogen-activated protein kinase kinase kinase), STE7 (mitogen-activated protein kinase kinase), and FUS3/KSS1 (mitogen-activated protein kinase) in response to signals from both Cdc42 and the heterotrimeric G proteins associated with transmembrane pheromone receptors. ACF5 is a human cDNA encoding a protein kinase homologous to STE20. This protein kinase, also designated HPK/GCK-like kinase (HGK), has nucleotide sequences that encode an open reading frame of 1165 amino acids with 11 kinase subdomains. HGK is a serine/threonine protein kinase that specifically activated the c-Jun N-terminal kinase (JNK) signaling pathway when transfected into 293T cells, but does not stimulate either the extracellular signal-regulated kinase or p38 kinase pathway. HGK also increased AP-1-mediated transcriptional activity in vivo. HGK may be a novel activator of the JNK pathway. The cascade may look like this: HGK -> TAK1 -> MKK4, MKK7 -> JNK kinase cascade, which may mediate the TNFalpha signaling pathway.

Paragraph beginning at line 4 of page 161 has been amended as follows:

ACF8 protein sequence (SEQ ID NO:127):

Gene name: Phospholipase A2, group IVC (cytosolic, calcium-independent)

Unigene number: Hs.18858

Probeset Accession #: AA054087 Protein Accession #: NP 003697.1

Pfam: none identified

Summary: ACF8 is a membrane-bound, calcium-independent PLA2, named cPLA2-gamma. The sequence encodes a 541-amino acid protein containing a domain with significant homology to the catalytic domain of the 85-kDa cPLA2 (cPLA2-alpha). cPLA2-gamma does not contain the regulatory calcium-dependent lipid binding (CaLB) domain found in cPLA2-alpha. cPLA2-gamma does contain two consensus motifs for lipid modification, a prenylation motif (-CCLA) at the C terminus and a myristoylation site at the N terminus. cPLA2-gamma demonstrates a preference for arachidonic acid at the sn-2 position of phosphatidylcholine as compared with palmitic acid. cPLA2-gamma encodes a 3-kilobase message, which is highly expressed in heart and skeletal muscle, suggesting a specific role in these tissues.

Application No.: 09/784,356

Page 76

Paragraph beginning at line 34 of page 161 has been amended as follows:

ACG1 protein sequence (SEQ ID NO:128):

Gene name: Carbohydrate (chondroitin 6/keratan) sulfotransferase 1

Unigene number: Hs.104576
Probeset Accession #: AA868063
Protein Accession #: NP\_003645.1

Pfam: none identified

Summary: Chondroitin 6-sulfotransferase (C6ST) is the key enzyme in the biosynthesis of chondroitin 6-sulfate, a glycosaminoglycan implicated in chondrogenesis, neoplasia, atherosclerosis, and other processes. C6ST catalyzes the transfer of sulfate from 3'-phosphoadenosine 5'-phosphosulfate to carbon 6 of the N-acetylgalactosamine residues of chondroitin.

Paragraph beginning at line 55 of page 161 has been amended as follows:

ACG5 protein sequence (SEQ ID NO:129):

Gene name: Multimerin
Unigene number: Hs.268107
Probeset Accession #: U27109
Protein Accession #: AAC52065

Signal sequence: prediction underlined

Pfam: EGF-like domain, C1q domain

Summary: Multimerin is a massive, soluble protein found in platelets and in the endothelium of blood vessels. Multimerin is composed of varying sized, disulfide-linked multimers, the smallest of which is a homotrimer. Multimerin is a factor V/Va-binding protein and may function as a carrier protein for platelet factor V. Northern analyses show a 4.7-kilobase transcript in cultured endothelial cells, a megakaryocytic cell line, platelets, and highly vascular tissues. The multimerin cDNA can encode a protein of 1228 amino acids with the probable signal peptide cleavage site between amino acids 19 and 20. The protein is predicted to be hydrophilic and to contain 23 N-glycosylation sites. The adhesive motif RGDS (Arg-Gly-Asp-Ser) and an epidermal growth factor-like domain were identified. Multimerin contains a probable coiled-coil structures in the central portion of its sequence. Additionally, the carboxyl-terminal region of multimerin resembles the globular, non-collagen-like, carboxyl-terminal domains of several other trimeric proteins, including complement Clq and collagens type VIII and X.

Paragraph beginning at line 32 of page 162 has been amended as follows:

ACC6 protein sequence (SEQ ID NO:131):

Gene name: Homo sapiens cDNA FLJ11502 fis, clone HEMBA1002102, weakly similar

to ANKRYIN

Unigene number: Hs.213194
Probeset Accession #: AA187101
Protein Accession #: none
Pfam: ankyrin repeats

Application No.: 09/784,356

Page 77

Paragraph beginning at line 45 of page 162 has been amended as follows:

ACC7 protein sequence (SEQ ID NO:132):

Gene name: Human RAL A gene Uniquene number: Hs.6906

Probeset Accession #: AA083572 cluster

Protein Accession #: P11233

Pfam: ras

Features: CAAX motif is underlined

Summary: The RALA gene encodes a low molecular mass ras-like GTP-binding protein that shares about 50% similarity with the ras proteins. GTP-binding proteins mediate the transmembrane signaling initiated by the occupancy of certain cell surface receptors. The RALA gene maps to 7p22-p15.

Paragraph beginning at line 63 of page 162 has been amended as follows:

ACC9 protein sequence (SEQ ID NO:133):

Gene name: KIAA0955 protein Uniquene number: Hs.10031

Probeset Accession #: AA027168 Protein Accession #: BAA76799.1

Pfam: CARD (Caspase recruitment domain)

Summary: Gene was originally isolated as a brain cDNA. The coding region contains a CARD domain, suggesting involvement in apoptotic signaling pathways.

Paragraph beginning at line 14 of page 163 has been amended as follows:

ACF6 Protein sequence (SEQ ID NO:134):

Gene name: Homo sapiens cDNA FLJ10669 fis, clone NT2RP2006275, weakly similar

to Microtubule-associated protein 1B [CONTAINS: LIGHT CHAIN LC1]

Unigene number: Hs.66048

Probeset Accession #: AA609717 Protein Accession #: BAA91743.1

Pfam: none identified

Summary: The cDNA for FLJ10669 was originally isolated from NT2 neuronal precursor cells (teratocarcinoma cell line) after 2-weeks of retinoic acid (RA) treatment. The protein sequence has similarity to microtubule-associated protein 1B (MAP-1B), suggesting a function for ACF6 in the regulating the cytoskeleton.

SF 1298767 v1